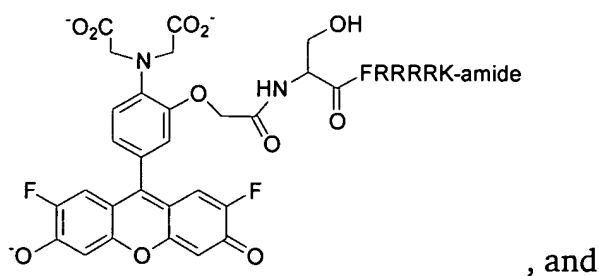
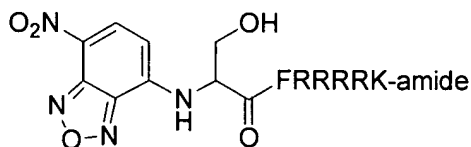


Amendments to the Claims:

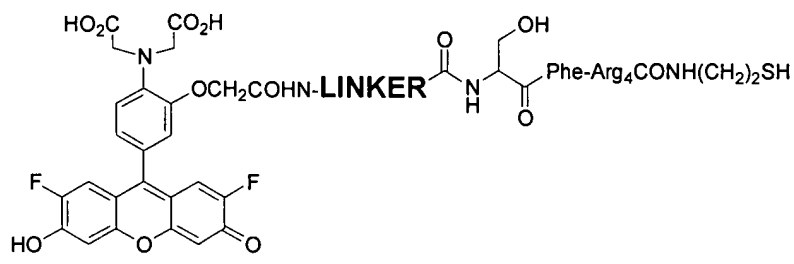
Please cancel Claims 103-121 and 144-145 without prejudice or disclaimer, and amend Claims 49, 56, 60-61, 88, 129 and 133 as set forth below.

1-48. (Canceled)

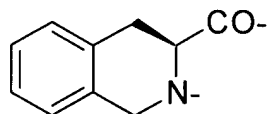
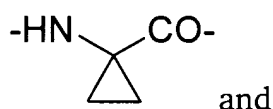
49. (Currently amended) A substrate for a protein kinase, ~~wherein the substrate comprises a peptide and at least one fluorophore, wherein a fluorophore is attached to a serine, a threonine, or a tyrosine on at least one terminal end of the peptide, wherein phosphorylation by the protein kinase of the terminal serine, the terminal threonine, or the terminal tyrosine to which the fluorophore is attached produces at least a 20% change in fluorescence intensity, and~~ wherein the substrate is selected from the group consisting of:



Applicant: David S. Lawrence
 Serial No.: 10/755,086
 Filed: January 9, 2004
 page 3 of 192



wherein F is phenylalanine, K is lysine, and R is arginine; and wherein the LINKER is selected from the group consisting of N-methyl glycine, L-proline, D-proline,



50-55. (Canceled)

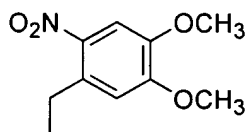
56. (Currently amended) A substrate for a protein kinase, wherein the substrate comprises:

a peptide substrate for the protein kinase, wherein the peptide comprises comprising a serine, a threonine, or a tyrosine on a terminal end of the peptide;

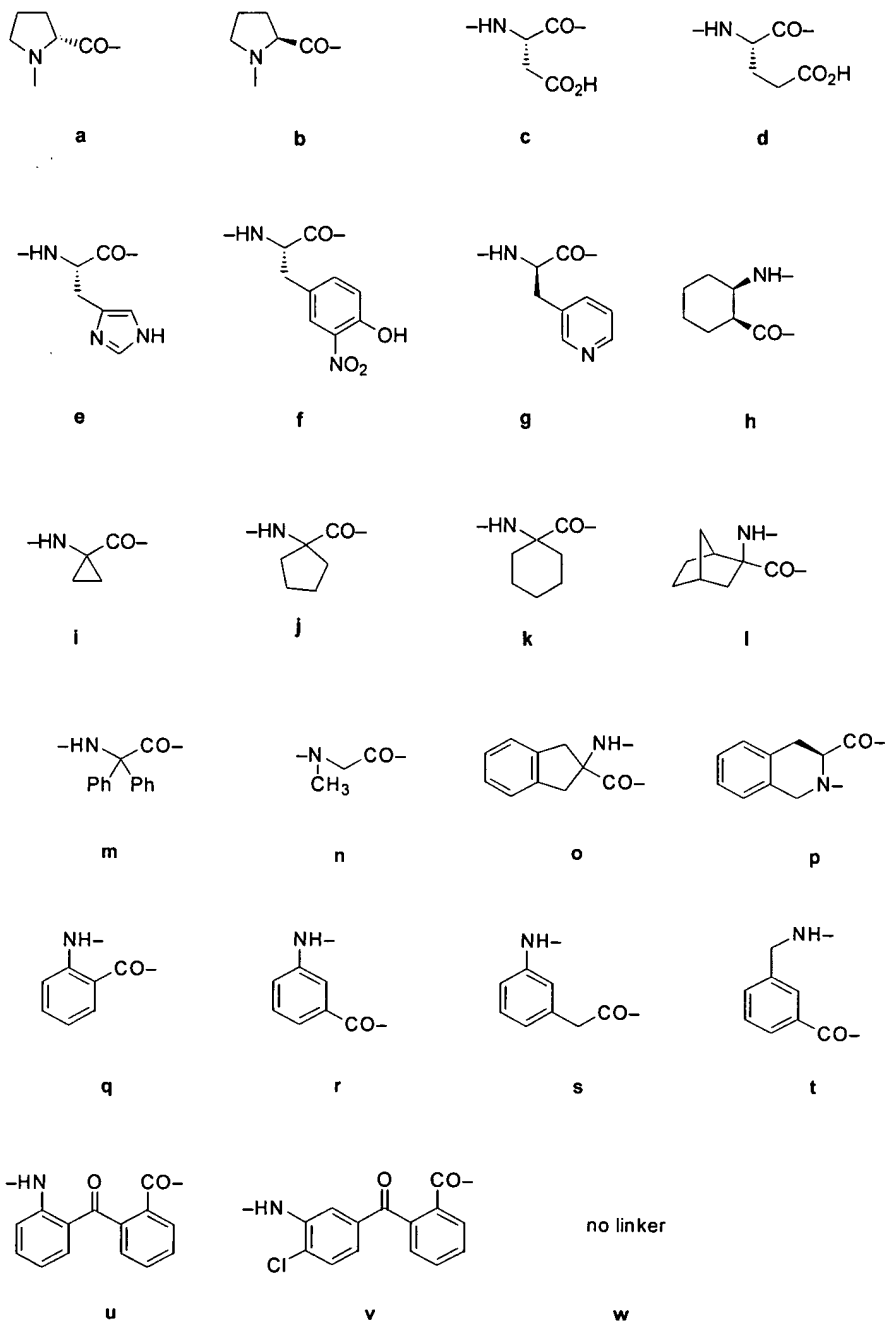
at least one fluorophore, wherein a fluorophore is attached to the serine, the threonine, or the tyrosine on the terminal end of the peptide; and

a photolabile side chain attached to the serine, the threonine, or the tyrosine on the terminal end of the peptide, wherein the photolabile side chain blocks transfer of a phosphoryl group from adenosine triphosphate to a hydroxyl moiety of the serine, the threonine, or the tyrosine so that the substrate cannot be phosphorylated by a protein kinase until the photolabile side chain is removed from the substrate; and

wherein the ~~the~~ photolabile side chain comprises the structure



or a fluorophore is attached to the peptide by a linker selected from the group consisting of



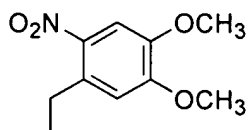
; and

wherein

- (i) the substrate is specific for a protein kinase subtype,
- (ii) the fluorophore is attached to the C-terminal end of the peptide,

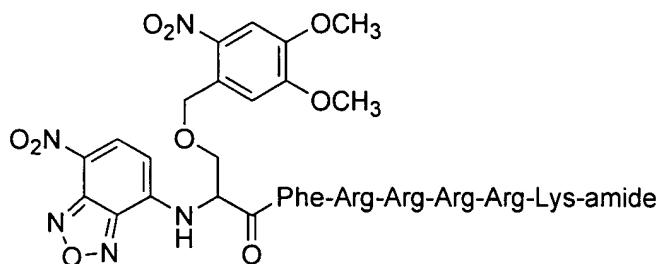
- (iii) a fluorophore is attached to each terminal end of the peptide,
- (iv) a first fluorophore is attached to a terminal end of the peptide and a second fluorophore, with photophysical properties distinct from the first fluorophore, is attached to any nonterminal site on the peptide,
- (v) the fluorophore is a 7-nitrobenz-2-oxa-1,3-diazole derivative,
- (vi) the fluorophore is attached to the peptide by a linker selected from the group consisting of a carboxamide linker, an aminobenzoic acid linker, a sulfonamide linker, a urea linker, a thiourea linker, an ester linker, a thioester linker, an alkylamine linker, an arylamine linker, an ether linker, and a thioether linker, and/or
- (vii) the substrate further comprises a carbohydrate, a lipid or a nucleic acid.

57. (Original) The substrate of claim 56, wherein the photolabile side chain comprises the structure



58. (Original) The substrate of claim 56, wherein the substrate comprises a serine with a photolabile side chain that blocks phosphoryl transfer.

59. (Original) The substrate of claim 58, wherein the substrate has the structure



60. (Original) The substrate of claim 56, wherein after removal of the photolabile side chain, phosphorylation by a protein kinase of the terminal serine, the terminal threonine, or the terminal tyrosine to which the fluorophore is attached produces at least a 20% change in fluorescence intensity.

61. (Currently amended) The substrate of claim 56, wherein after removal of the photolabile side chain, phosphorylation by a protein kinase of the terminal serine, the terminal threonine, or the terminal tyrosine to which the fluorophore is attached produces at least a 20% increase in fluorescence intensity. ~~The substrate of claim 60, wherein the change in fluorescence intensity when the substrate is phosphorylated by the protein kinase is an increase in fluorescence intensity.~~

62. (Currently amended) The substrate of claim 56, wherein after removal of the photolabile side chain, phosphorylation by a protein kinase of the terminal serine, the terminal threonine, or the terminal tyrosine to which the fluorophore is attached produces at least a 20% decrease in fluorescence intensity. ~~The substrate of claim 60, wherein the change in fluorescence intensity when the substrate is phosphorylated by the protein kinase is a decrease in fluorescence intensity.~~

63. (Previously presented) The substrate of claim 60, wherein phosphorylation of

the substrate by the protein kinase produces at least a 70% change in fluorescence intensity.

64. (Original) The substrate of claim 63, wherein phosphorylation of the substrate by the protein kinase produces at least a 100% change in fluorescence intensity.

65. (Original) The substrate of claim 64, wherein phosphorylation of the substrate by the protein kinase produces at least a 150% change in fluorescence intensity.

66. (Original) The substrate of claim 65, wherein phosphorylation of the substrate by the protein kinase produces at least a 250% change in fluorescence intensity.

67. (Previously presented) The substrate of claim 56, wherein the substrate is specific for a protein kinase subtype.

68. (Original) The substrate of claim 67, wherein the substrate is specific for protein kinase C.

69. (Original) The substrate of claim 68, wherein the substrate is specific for isoforms α , β , and γ of protein kinase C.

70. (Withdrawn) The substrate of claim 67, wherein the substrate is specific for protein kinase A, protein kinase B, protein kinase D, protein kinase G, Ca^{+} /calmodulin-dependent protein kinase, mitogen-activated protein kinase, protein kinase mos, protein kinase raf, protein tyrosine kinase, tyrosine kinase abl, tyrosine kinase src, tyrosine kinase yes, tyrosine kinase fps, tyrosine kinase met, cyclin-dependent protein kinase, or cdc2

kinase.

71. (Previously presented) The substrate of claim 56, wherein the substrate further comprises a carbohydrate, a lipid or a nucleic acid.

72. (Canceled)

73. (Previously presented) The substrate of claim 56, wherein the fluorophore is attached to the C-terminal end of the peptide.

74. (Previously presented) The substrate of claim 56, wherein the fluorophore is attached to the N-terminal end of the peptide.

75. (Previously presented) The substrate of claim 56, wherein a fluorophore is attached to each terminal end of the peptide.

76. (Original) The substrate of claim 75, wherein fluorophores with distinct photophysical properties are attached to different terminal ends of the peptide.

77. (Previously presented) The substrate of claim 56, wherein a first fluorophore is attached to a terminal end of the peptide and a second fluorophore, with photophysical properties distinct from the first fluorophore, is attached to any nonterminal site on the peptide.

78. (Previously presented) The substrate of claim 56, wherein the fluorophore is a 7-nitrobenz-2-oxa-1,3-diazole derivative.

79. (Withdrawn) The substrate of claim 56, wherein the fluorophore is a fluorescein derivative.

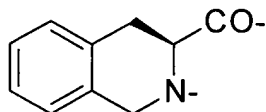
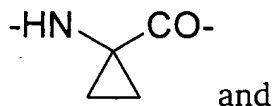
80. (Withdrawn) The substrate of claim 56, wherein the fluorophore is selected from the group consisting of a dansyl derivative, an acridine derivative, an Alexa Fluor derivative, a BODIPY derivative, an Oregon Green derivative, a Rhodamine Green derivative, a Rhodamine Red-X derivative, a Texas Red derivative, a Cascade Blue derivative, a Cascade Yellow derivative, a Marina Blue derivative, a Pacific Blue derivative, an AMCA-X derivative, and a coumarin derivative.

81. (Canceled)

82. (Withdrawn) The substrate of claim 56, wherein the fluorophore is attached to the peptide by a metal chelating linker.

83. (Previously presented) The substrate of claim 56, wherein the fluorophore is attached to the peptide by a linker selected from the group consisting of a carboxamide linker, an aminobenzoic acid linker, a sulfonamide linker, a urea linker, a thiourea linker, an ester linker, a thioester linker, an alkylamine linker, an arylamine linker, an ether linker, and a thioether linker.

84. (Withdrawn) The substrate of claim 56, wherein the fluorophore is attached to the peptide by a linker selected from the group consisting of N-methyl glycine, L-proline, D-proline,

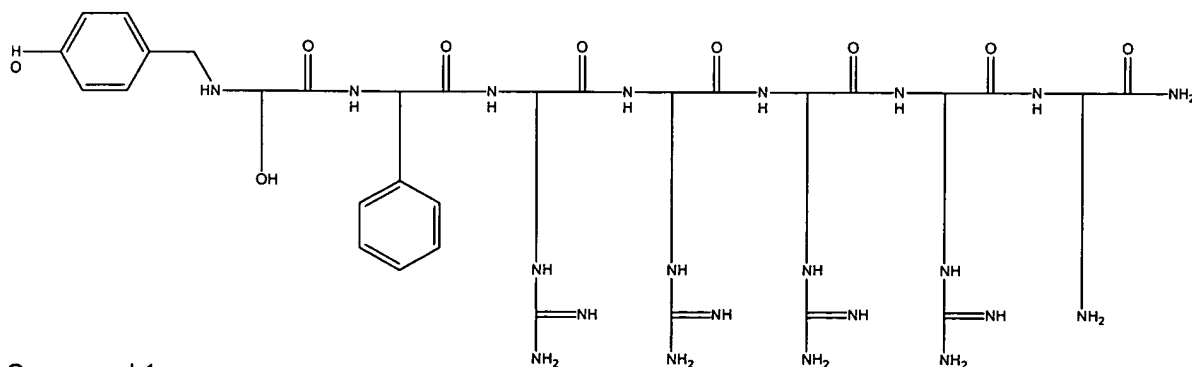


85. (Canceled)

86. (Previously presented) A composition comprising the substrate of claim 56, and a carrier.

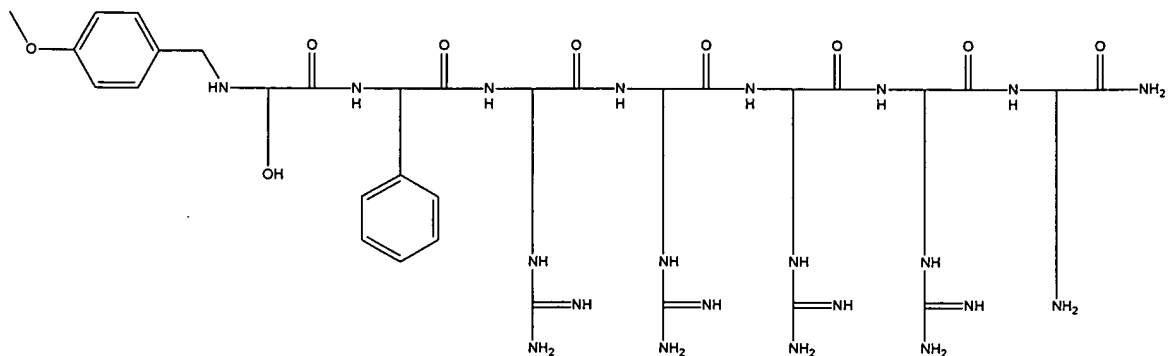
87. (Original) The composition of claim 86, wherein the composition is a pharmaceutical composition and the carrier is a pharmaceutically acceptable carrier.

88. (Currently amended) A chemical compound selected from the group of compounds consisting of: ~~set forth in Table 3.~~

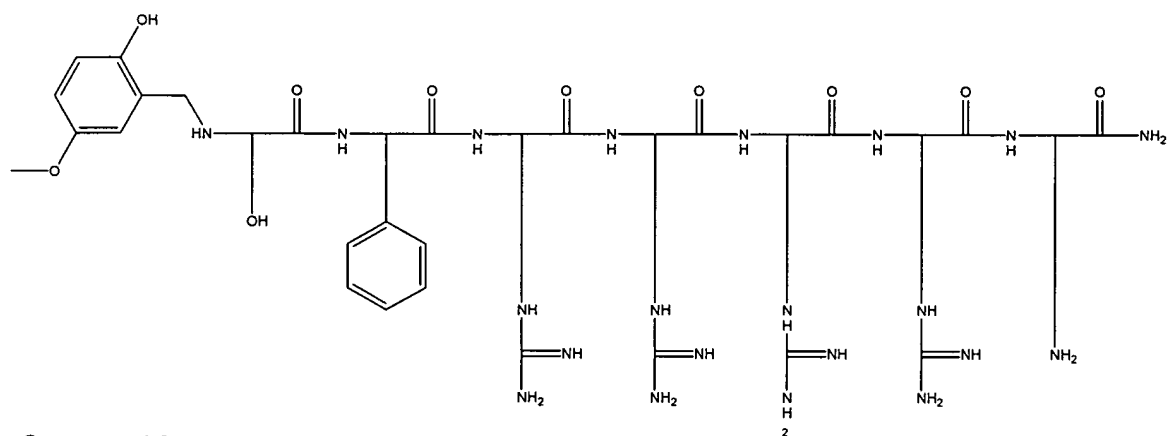


Compound 1

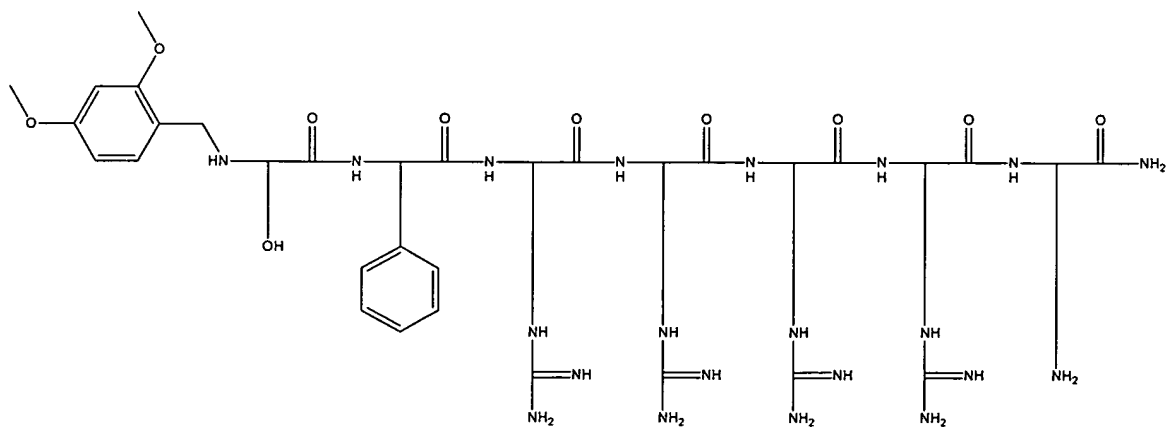
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 12 of 192



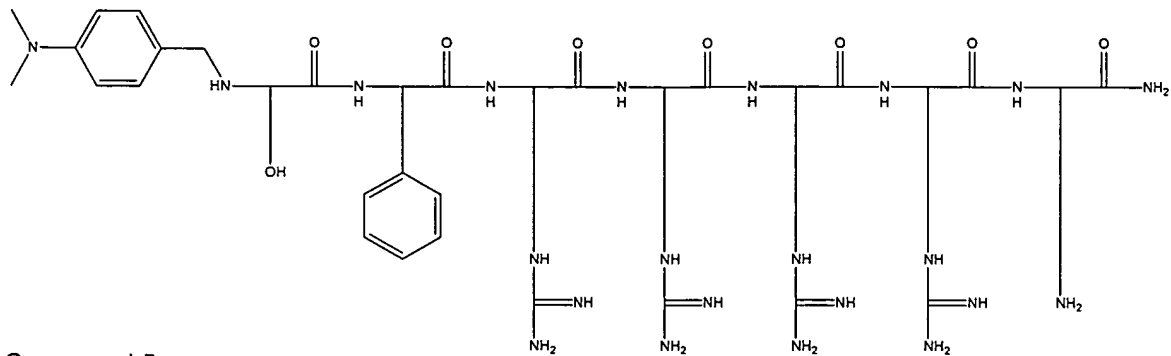
Compound 2



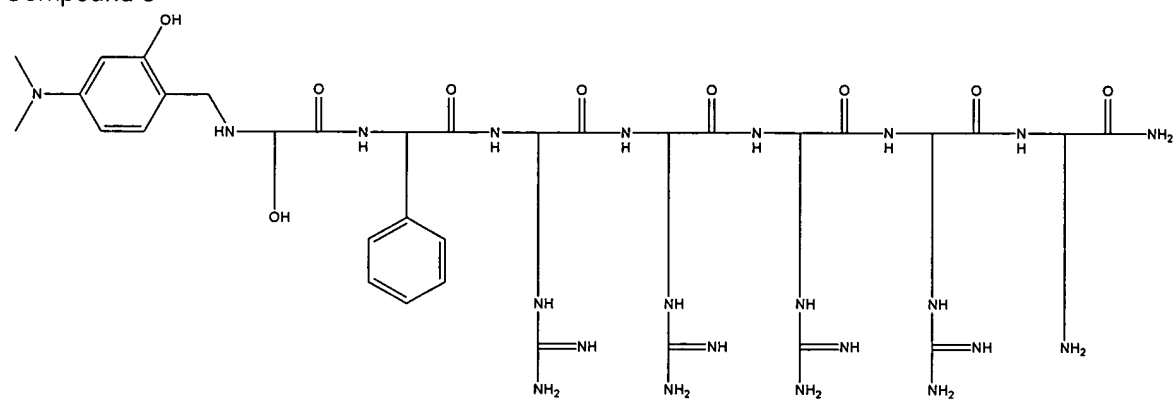
Compound 3



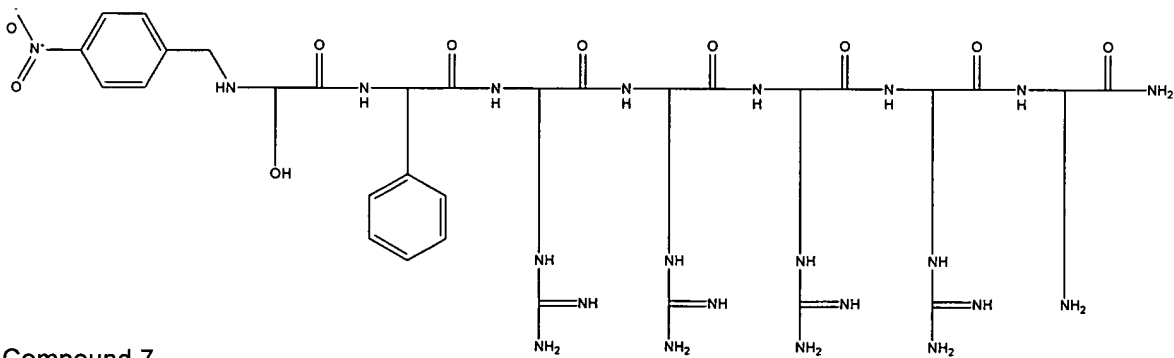
Compound 4



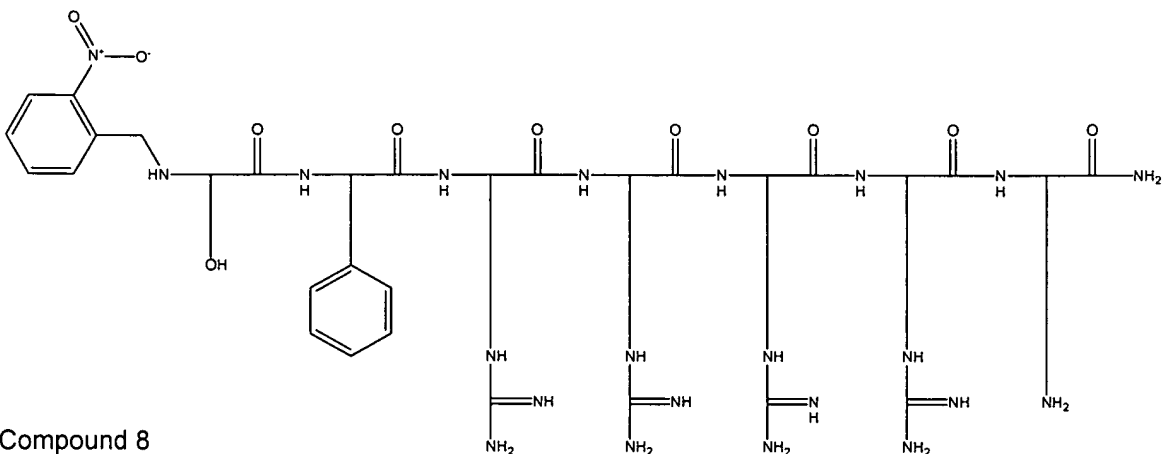
Compound 5



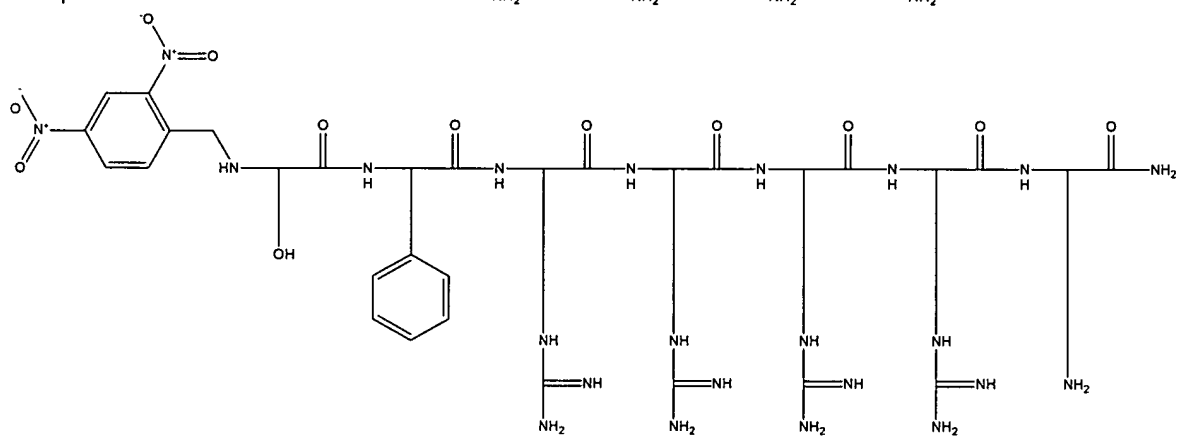
Compound 6



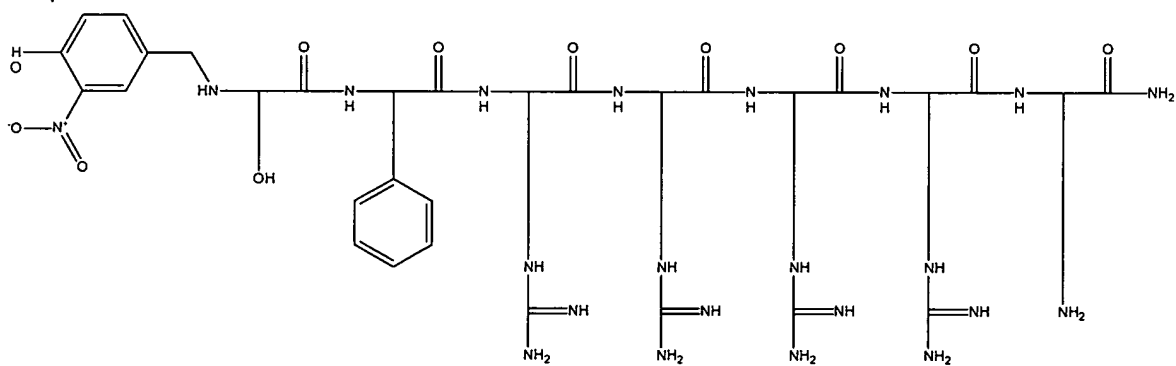
Compound 7



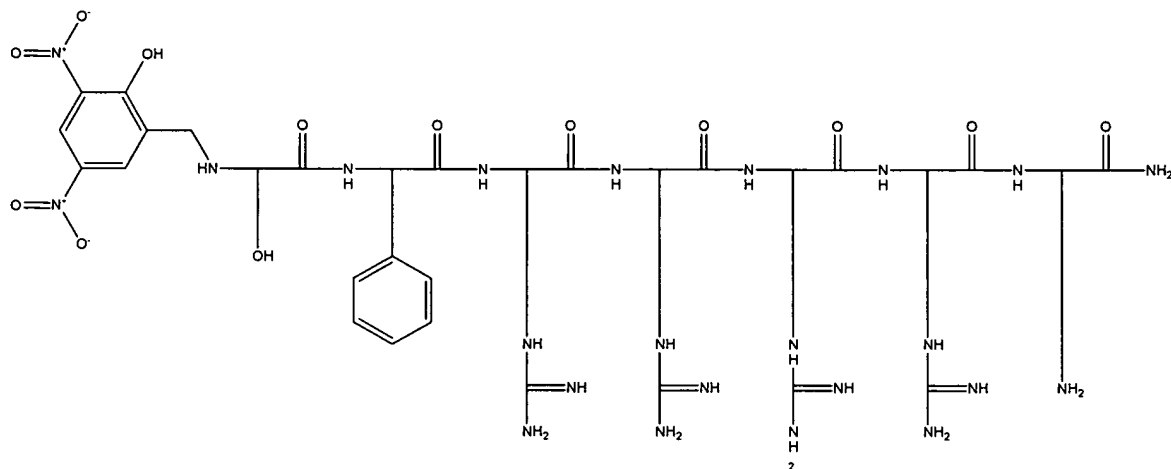
Compound 8



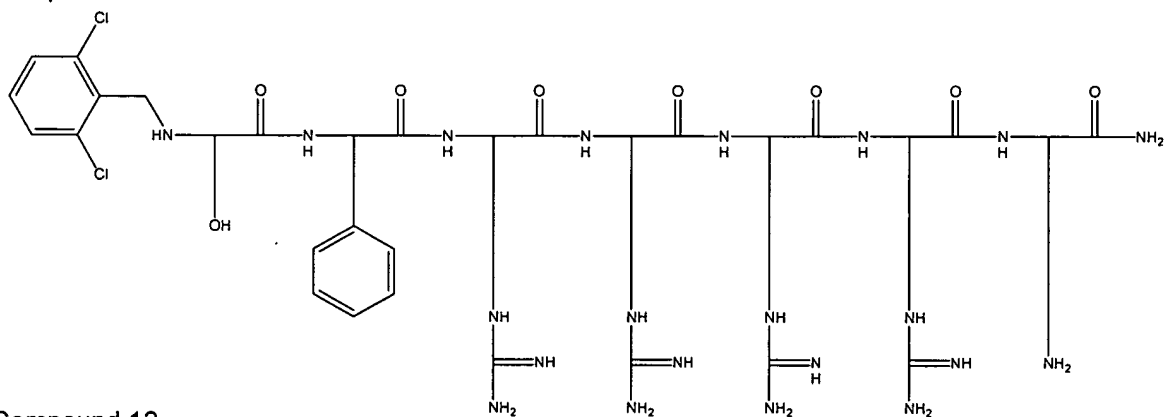
Compound 9



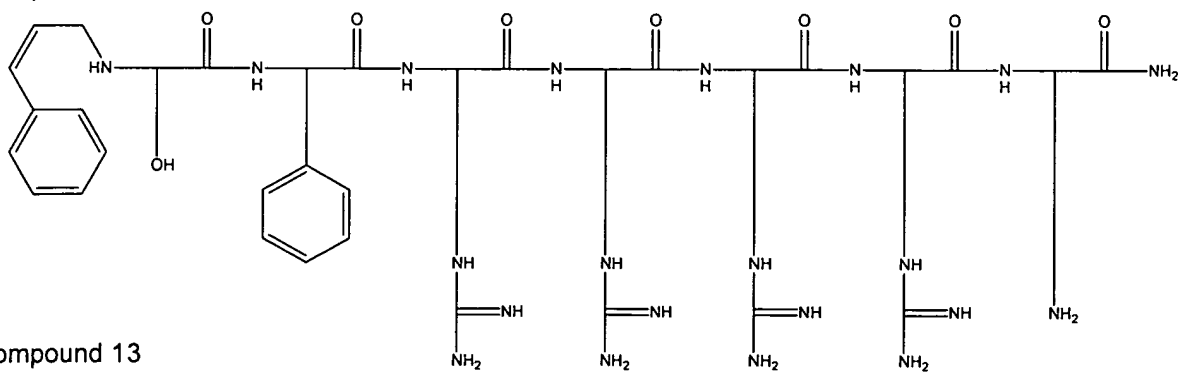
Compound 10



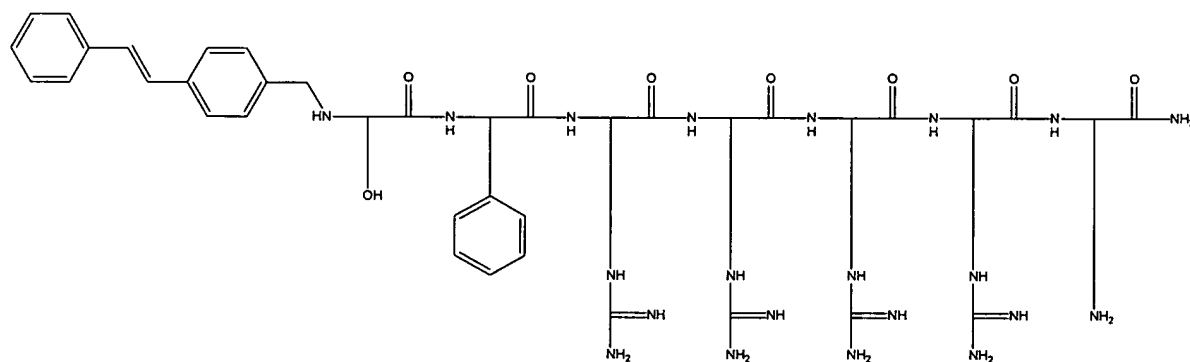
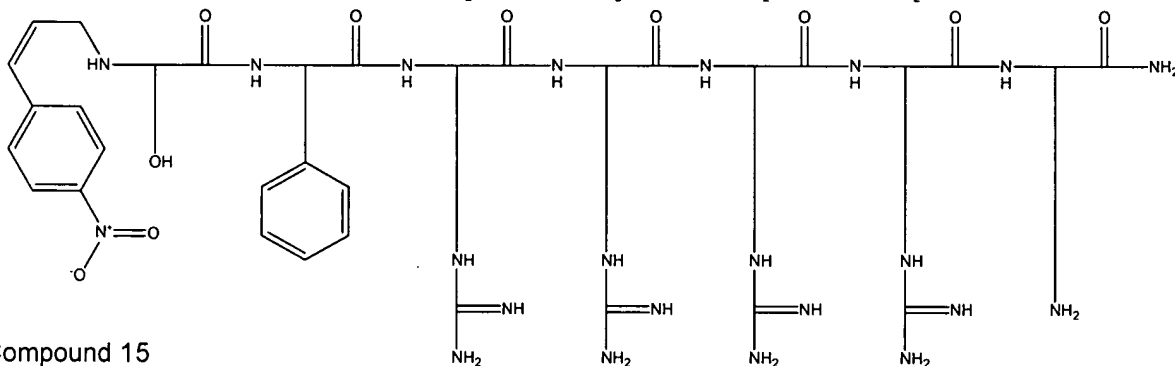
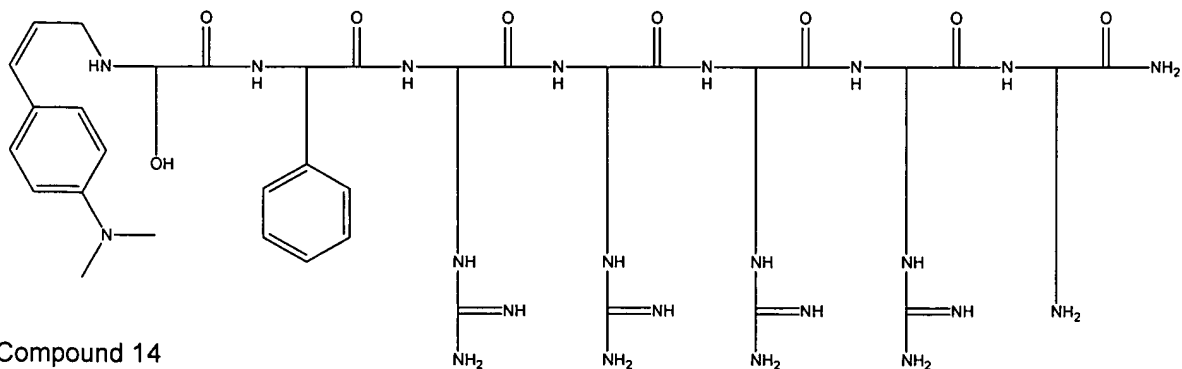
Compound 11

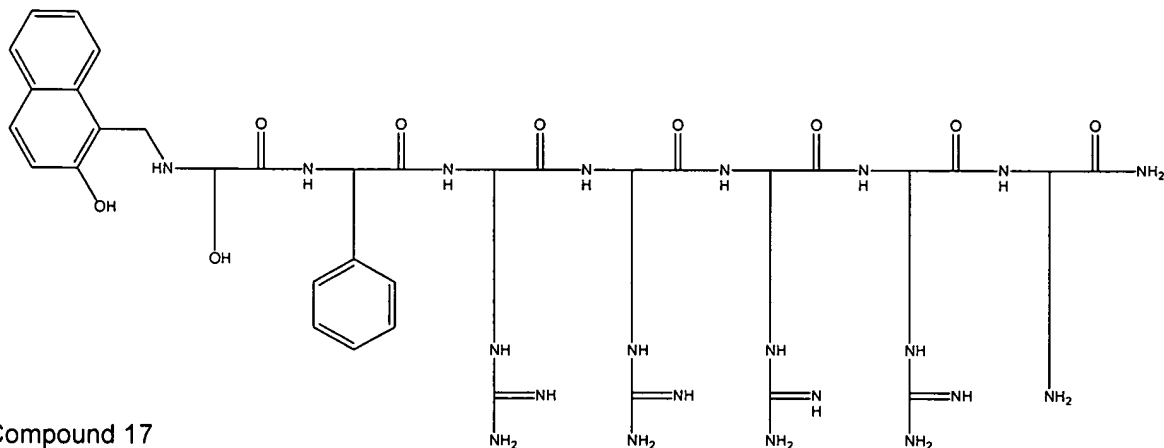


Compound 12

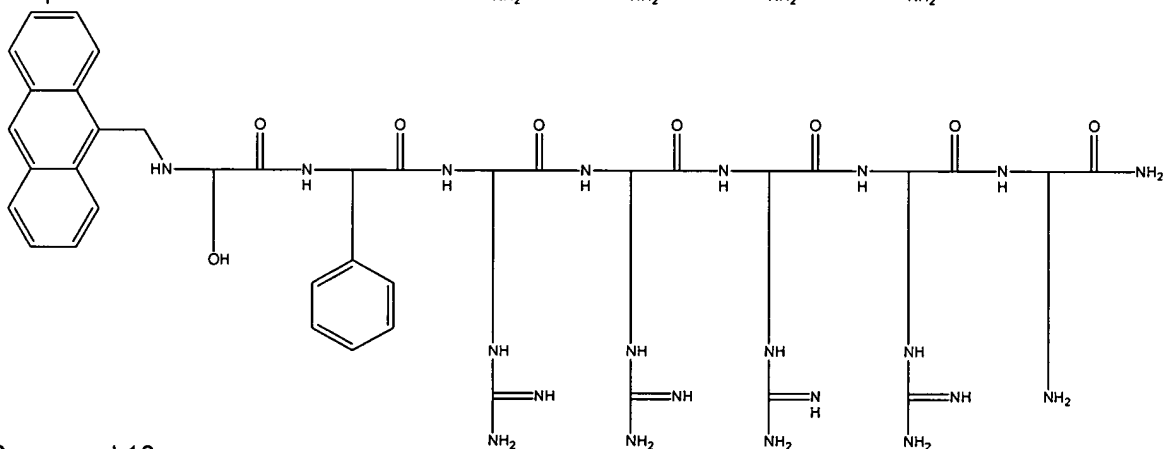


Compound 13

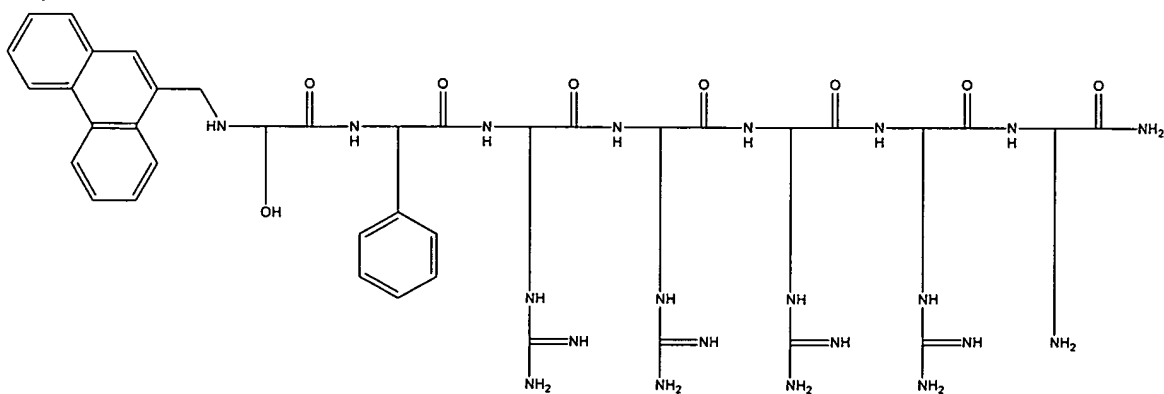




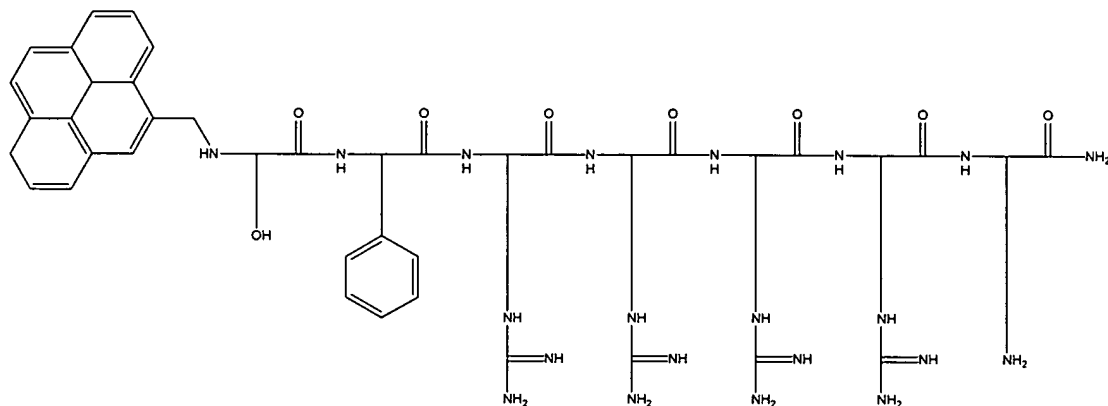
Compound 17



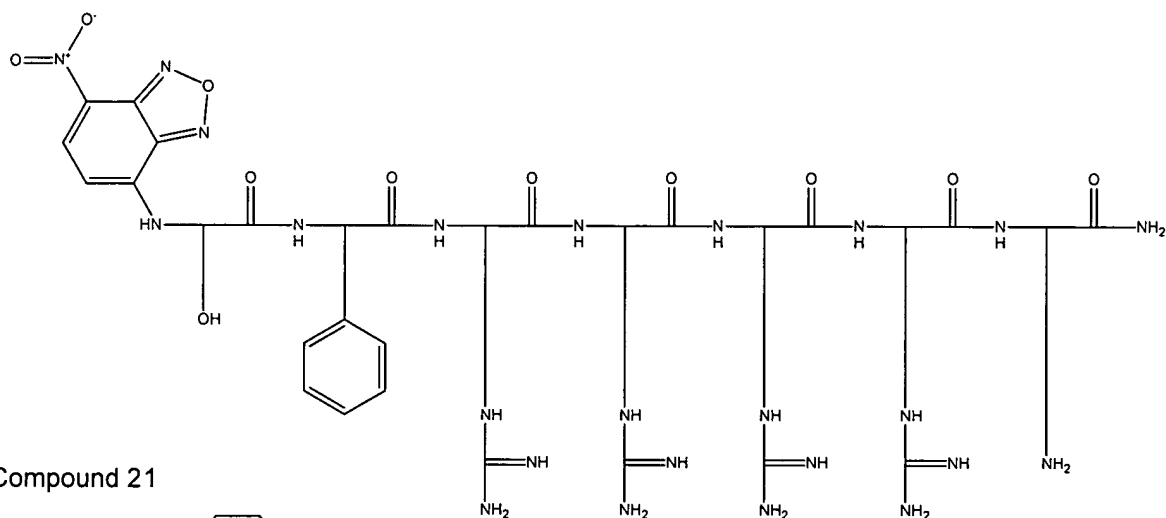
Compound 18



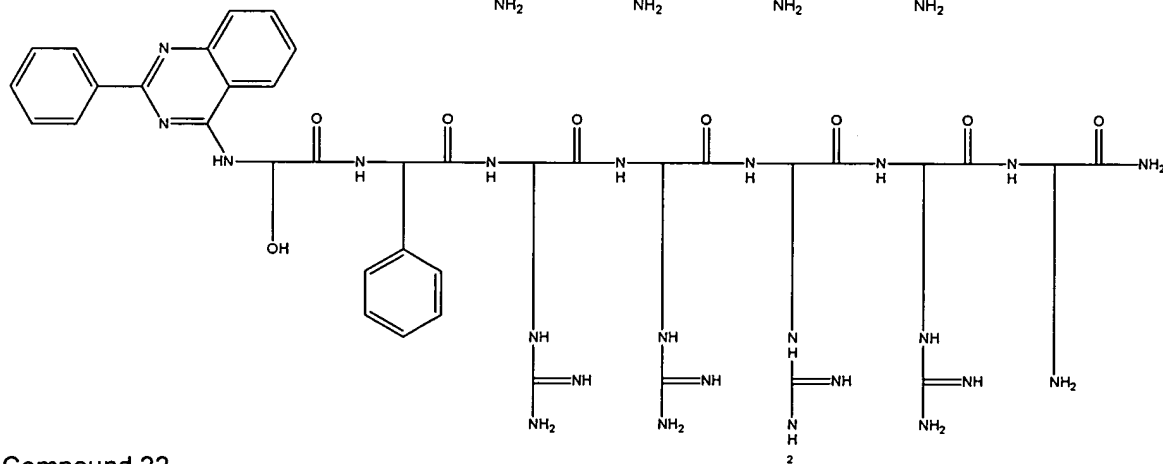
Compound 19



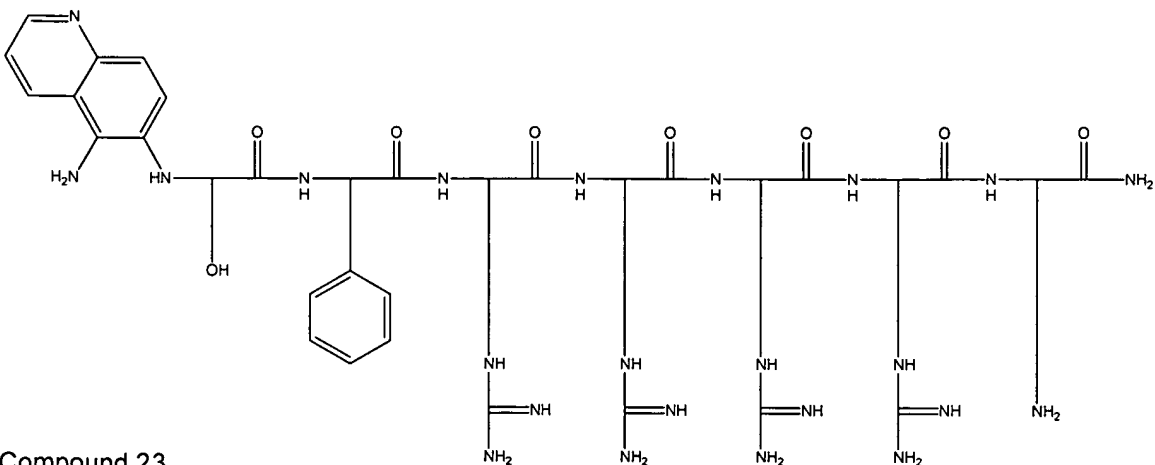
Compound 20



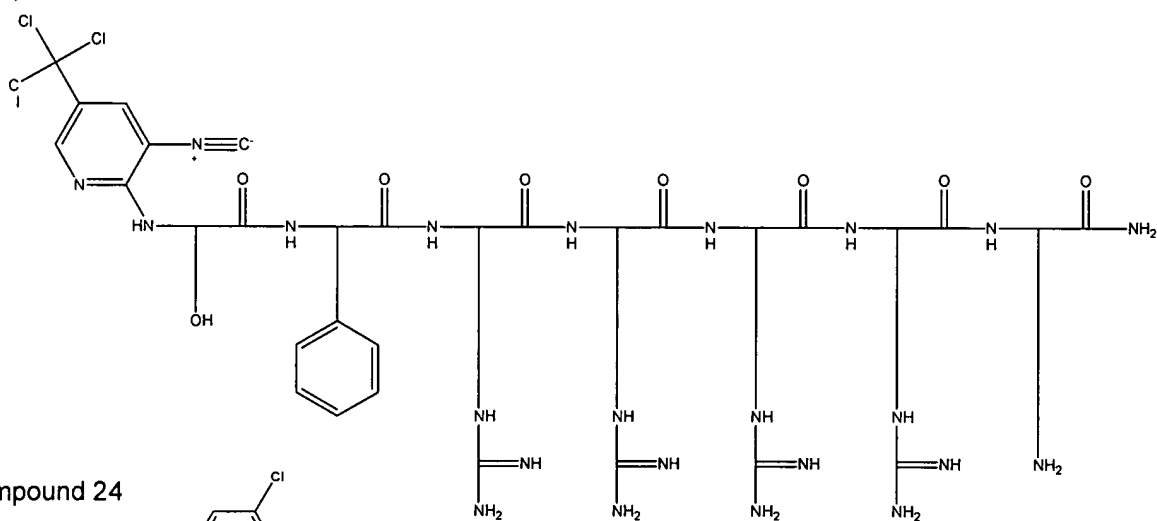
Compound 21



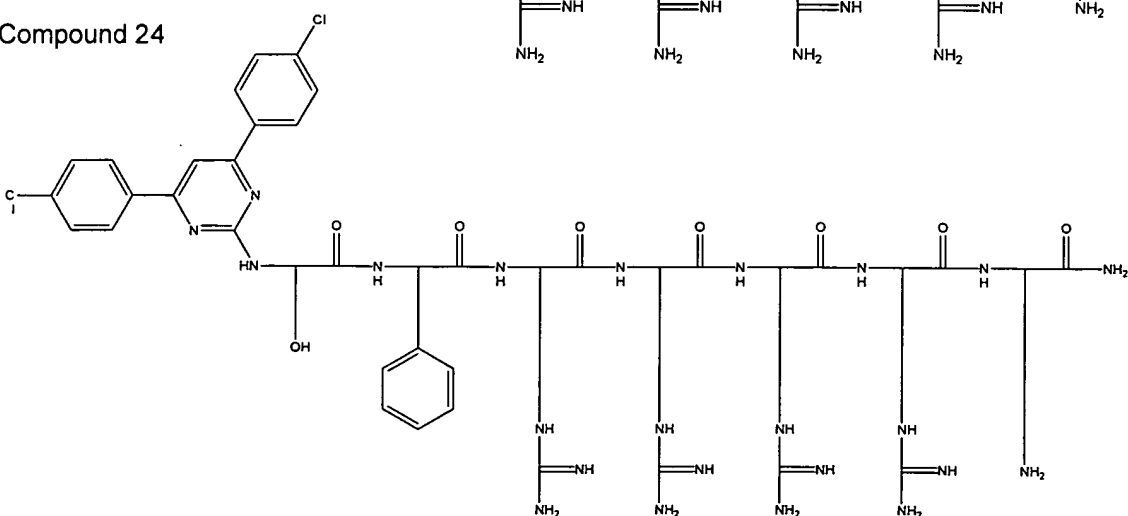
Compound 22



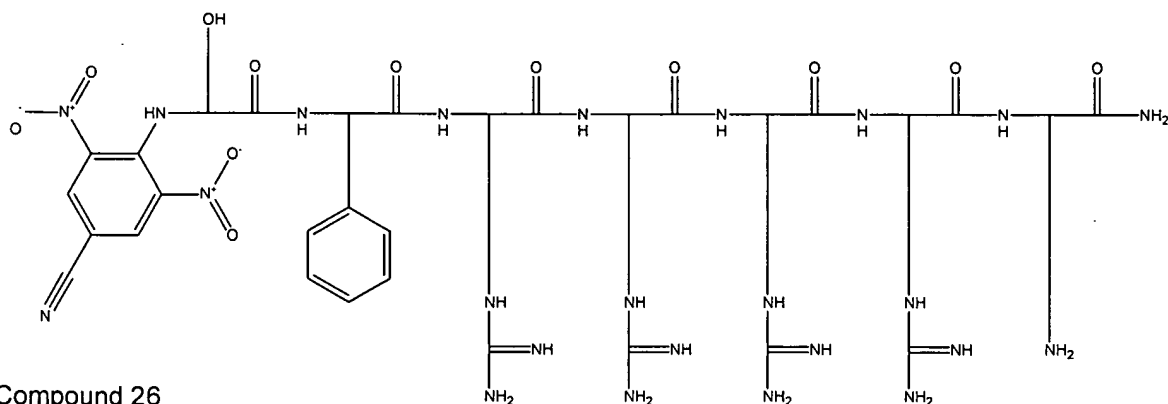
Compound 23



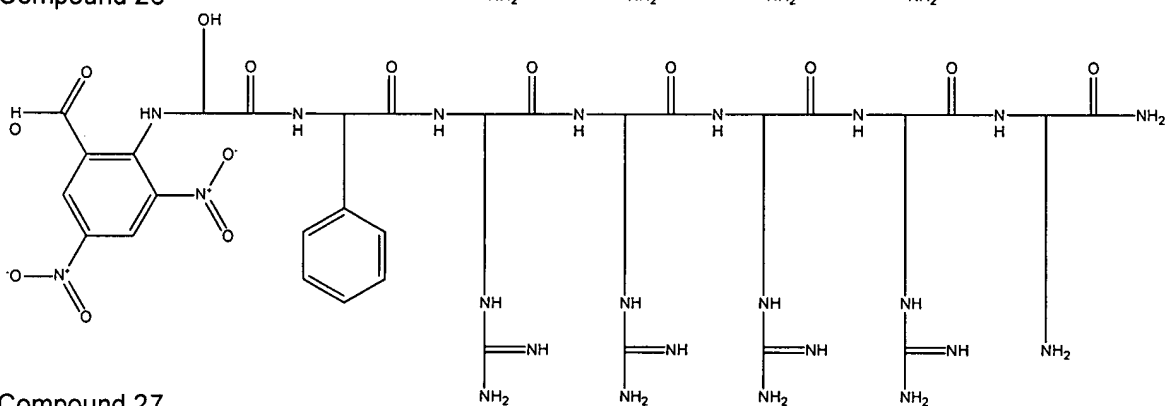
Compound 24



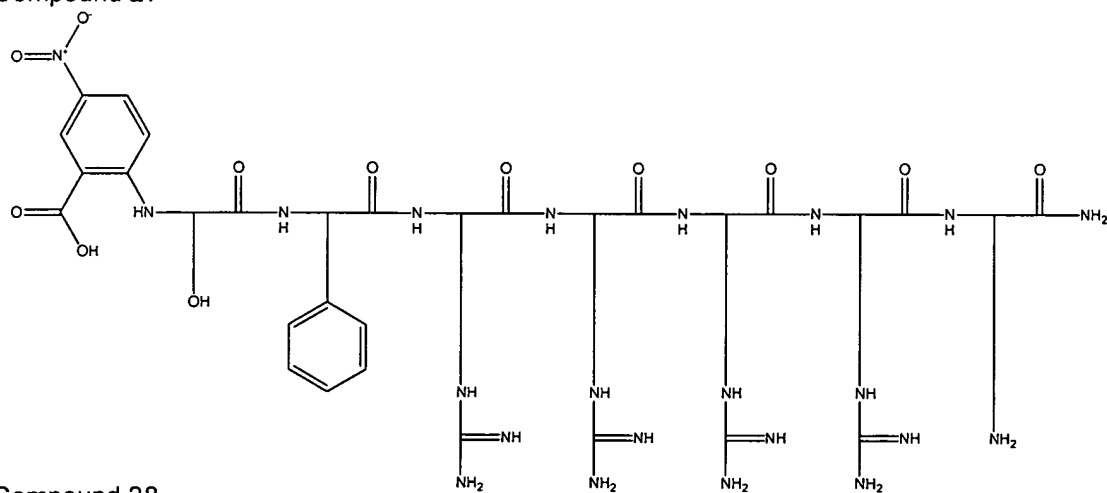
Compound 25



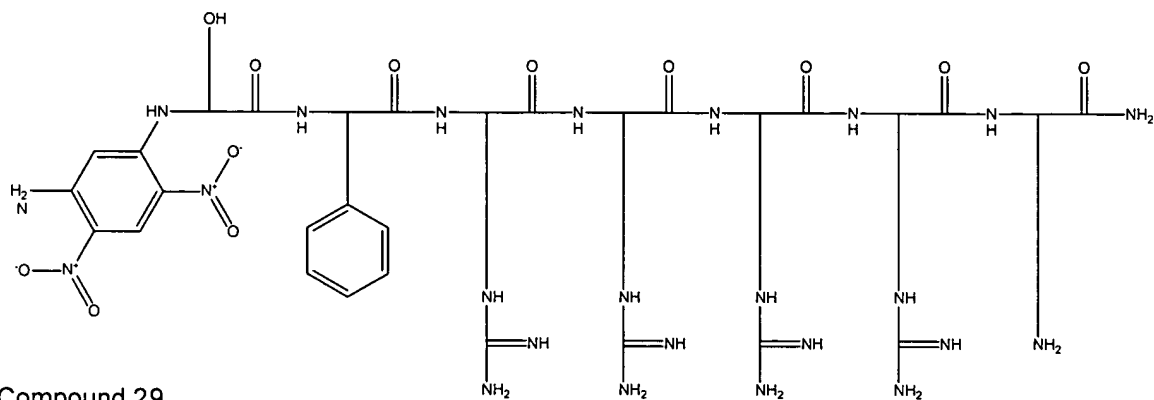
Compound 26



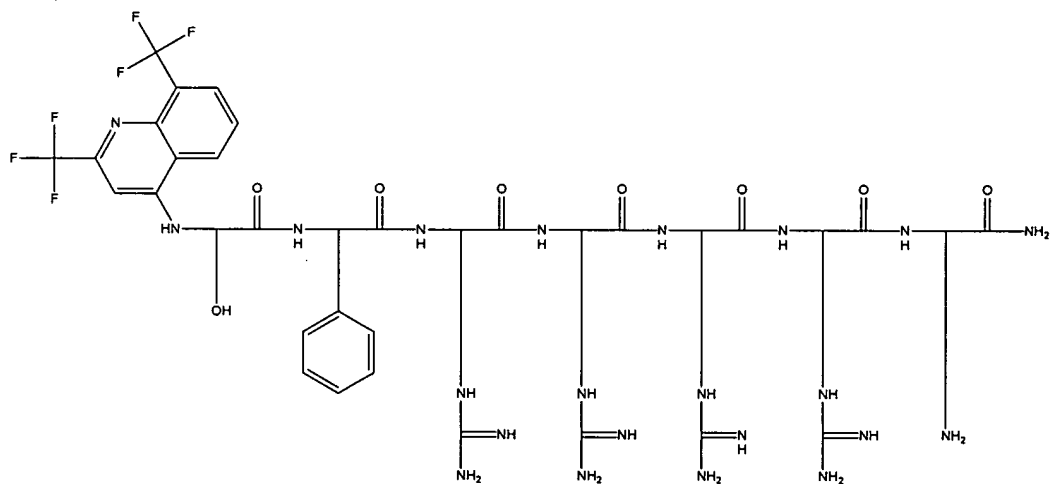
Compound 27



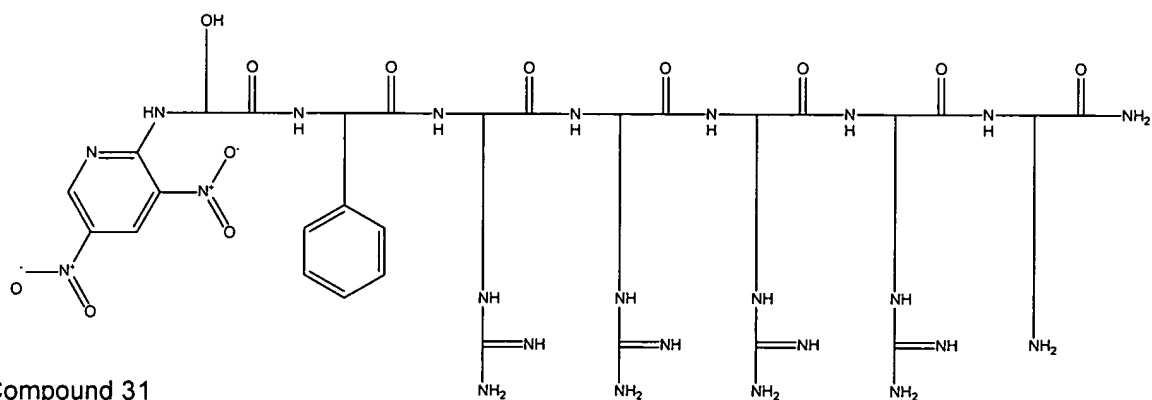
Compound 28



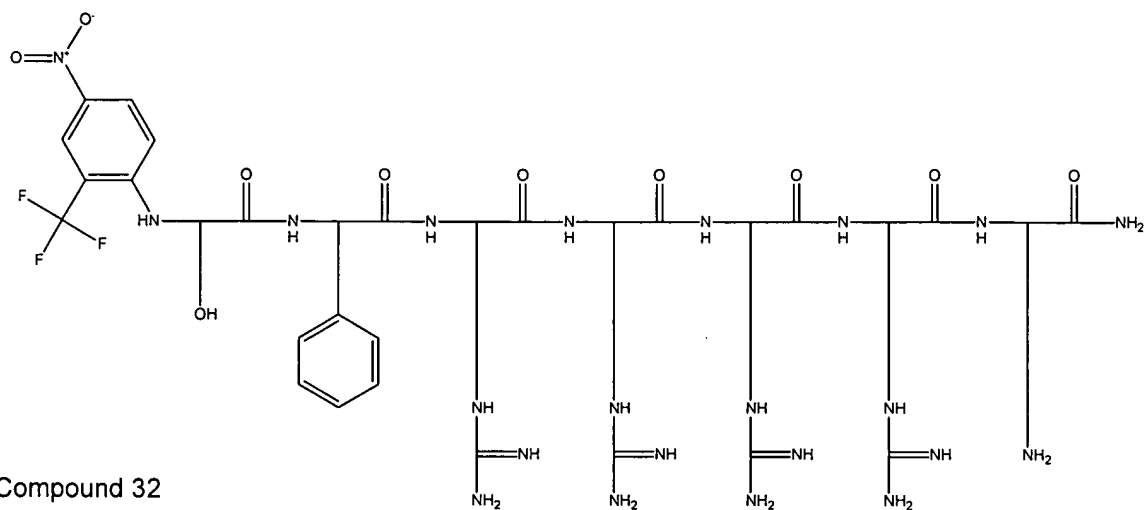
Compound 29



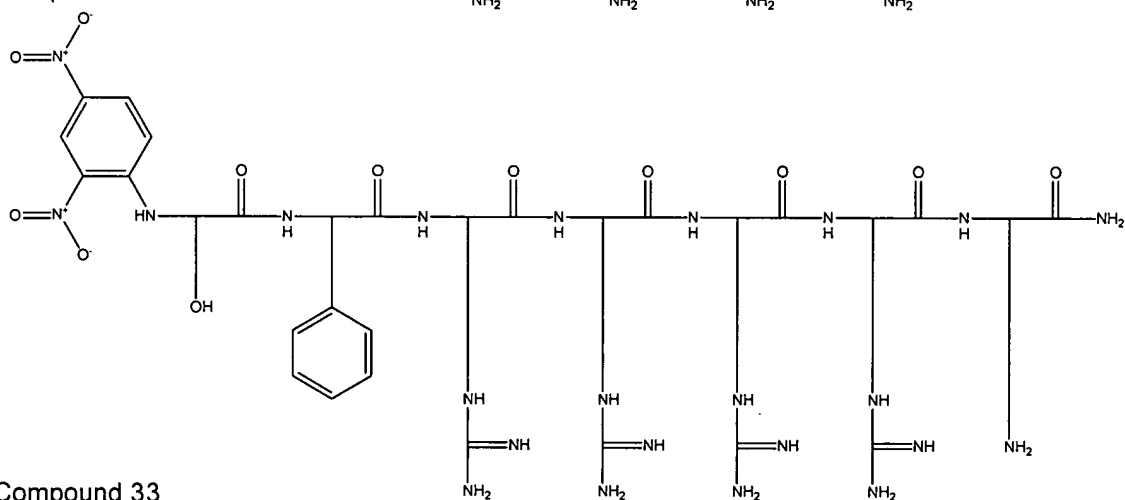
Compound 30



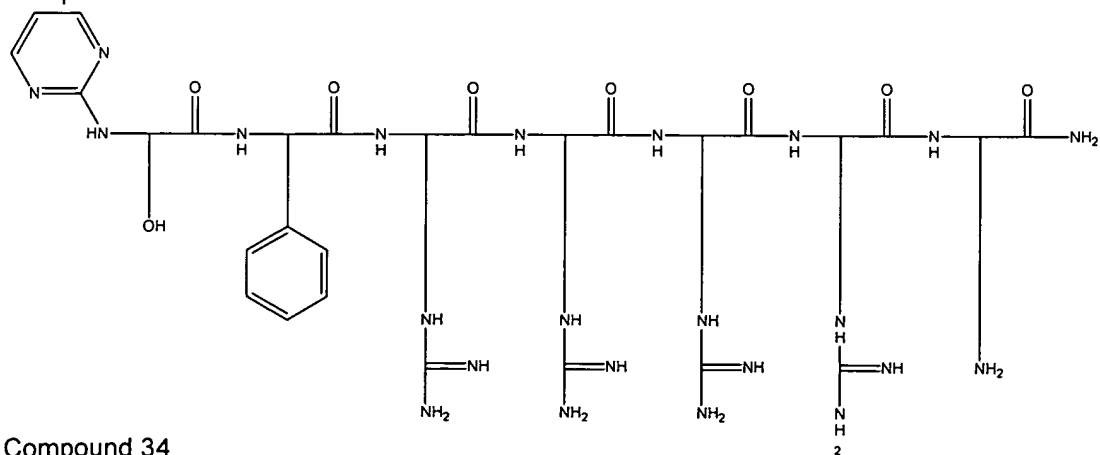
Compound 31



Compound 32

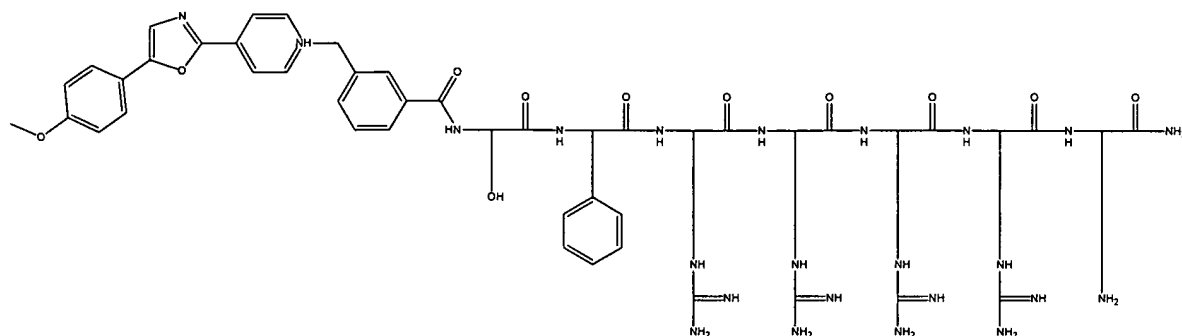


Compound 33

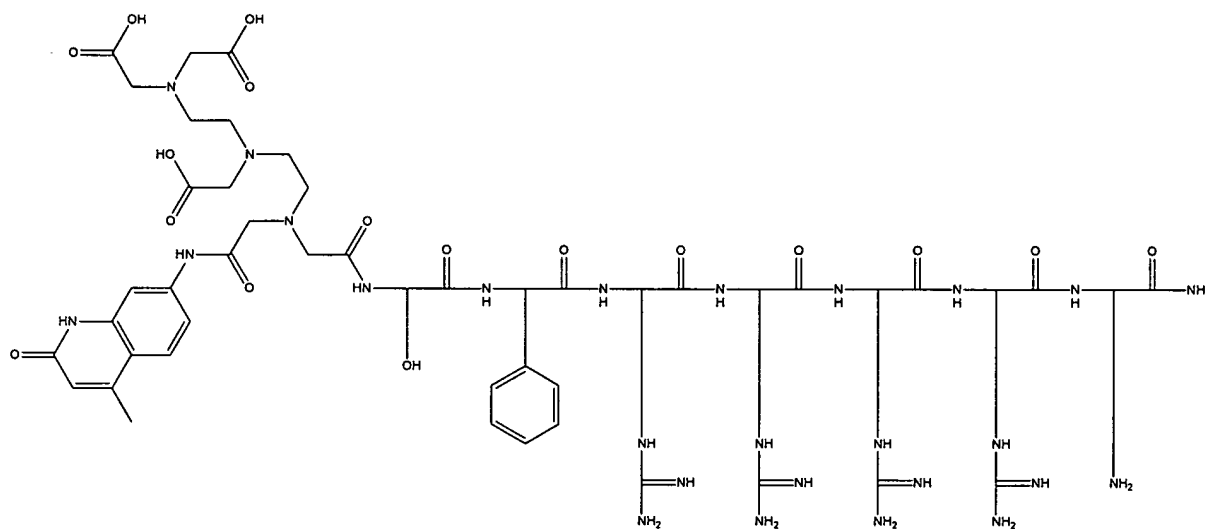


Compound 34

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 23 of 192

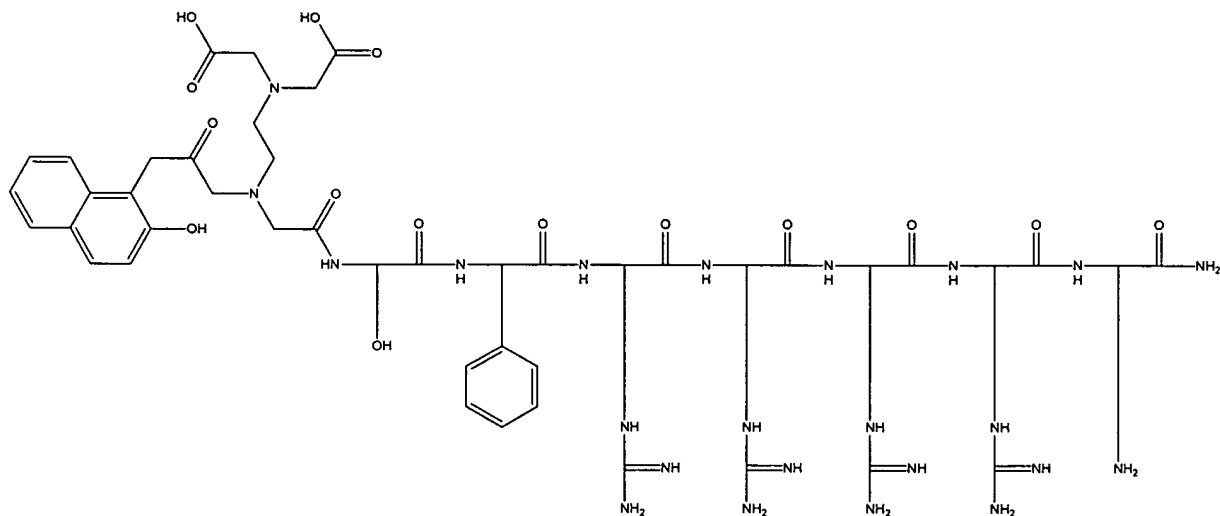


Compound 35

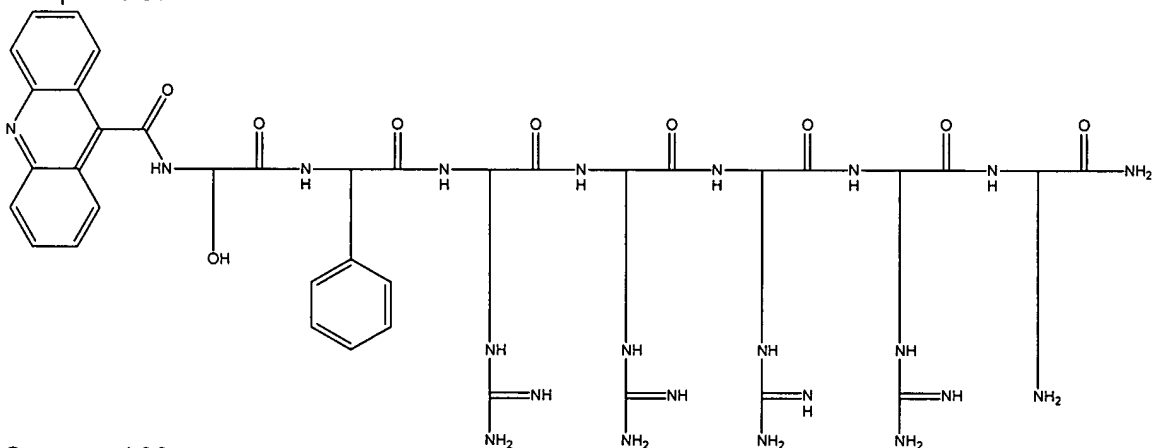


Compound 36

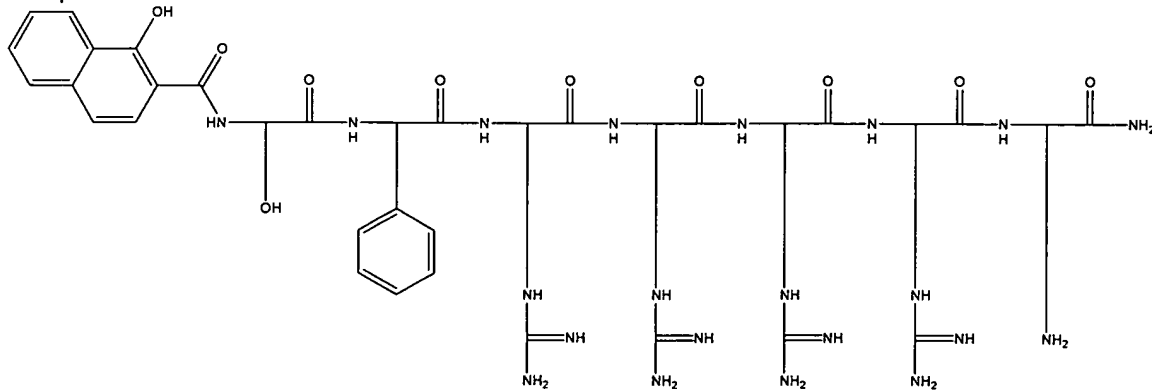
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 24 of 192



Compound 37

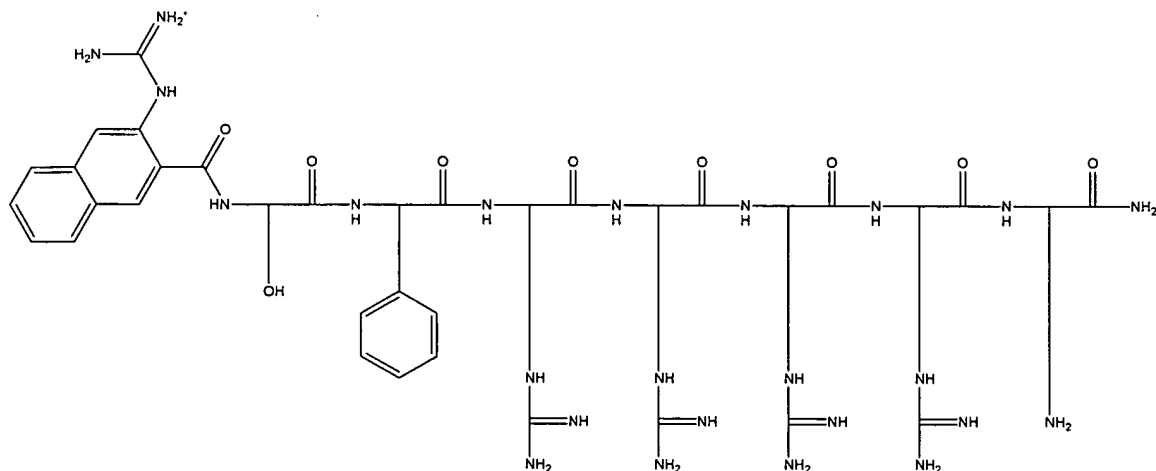


Compound 38

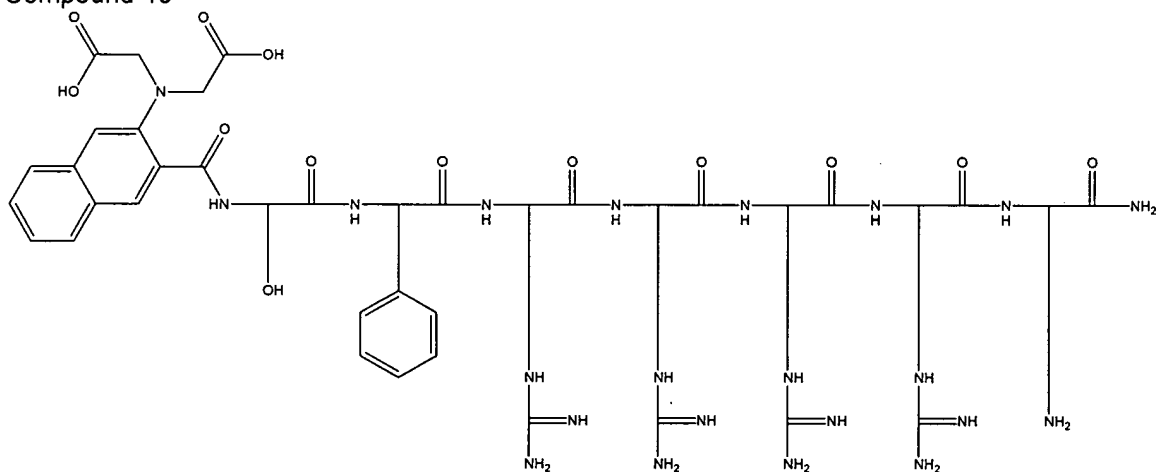


Compound 39

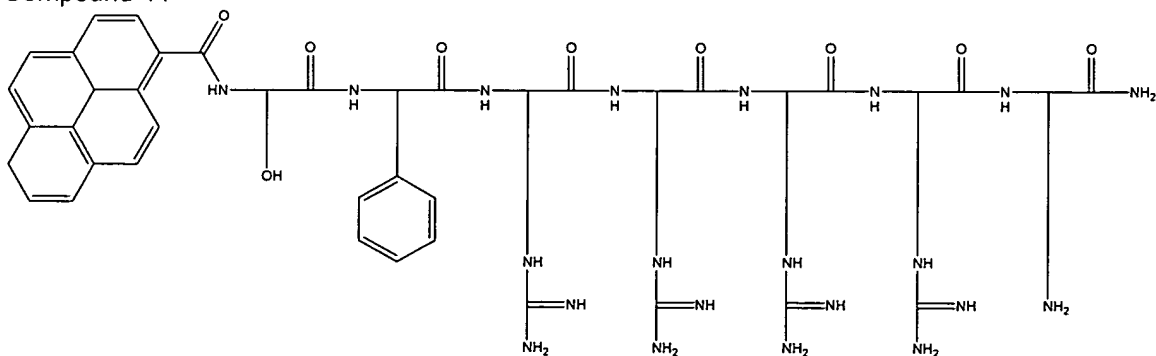
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 25 of 192



Compound 40



Compound 41



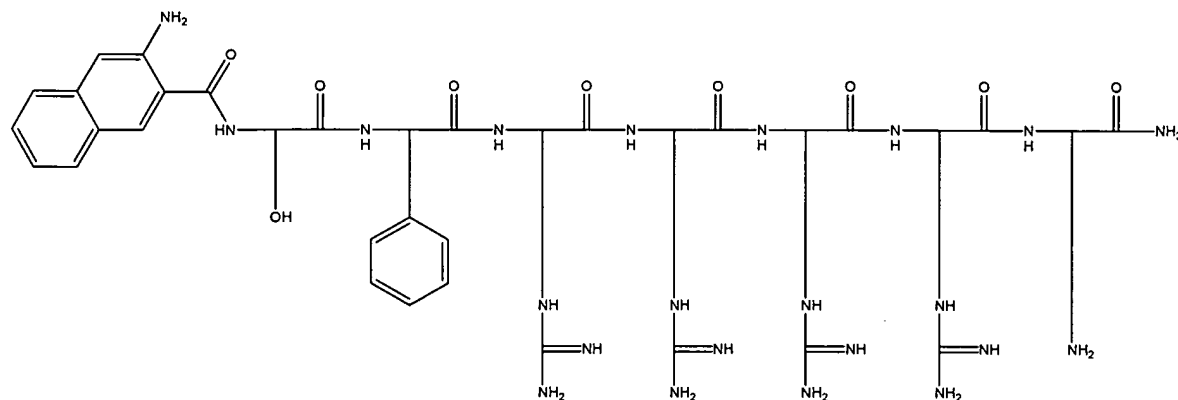
Compound 42

Applicant: David S. Lawrence

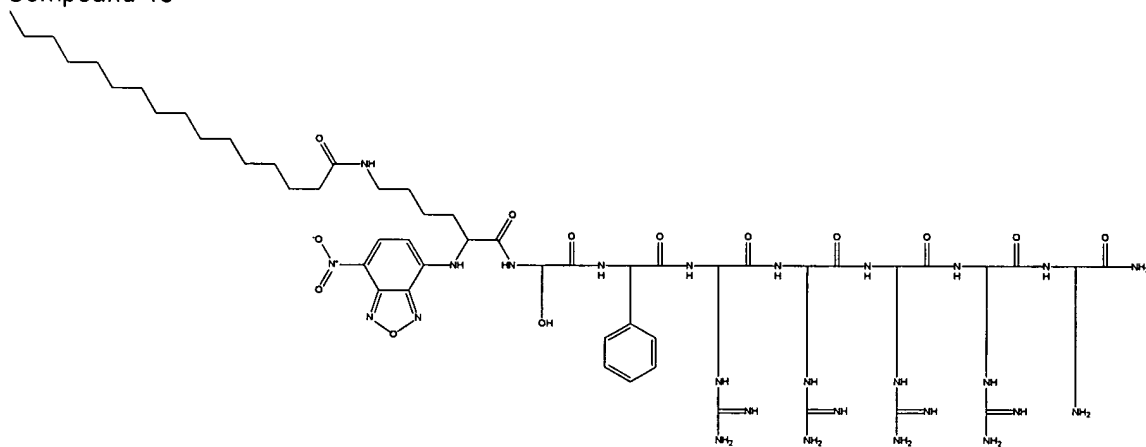
Serial No.: 10/755,086

Filed: January 9, 2004

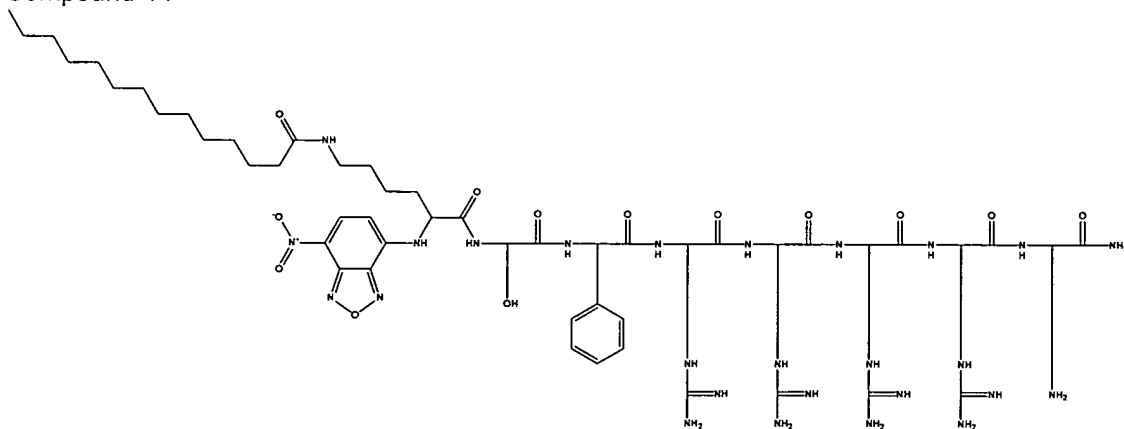
page 26 of 192



Compound 43

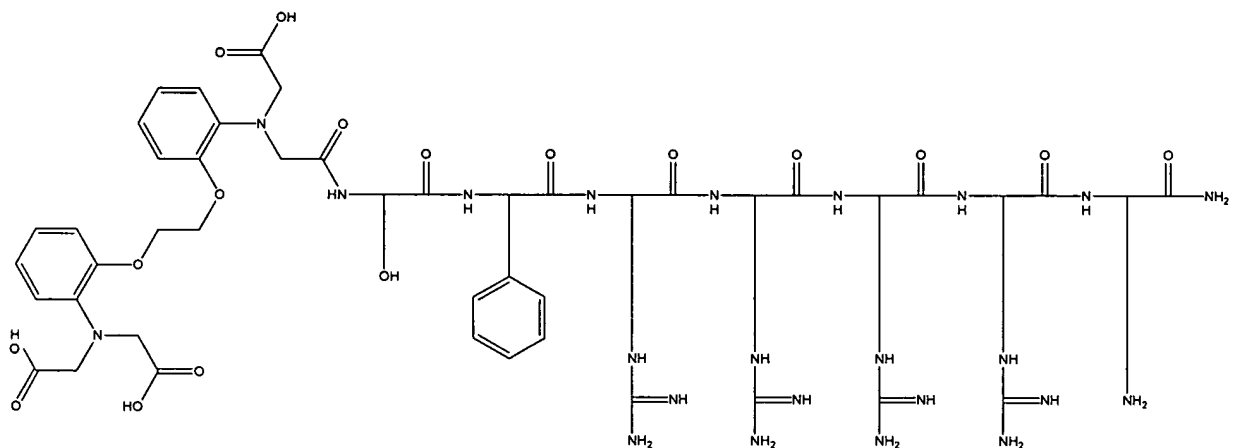


Compound 44



Compound 45

page 27 of 192

[illegible][illegible]

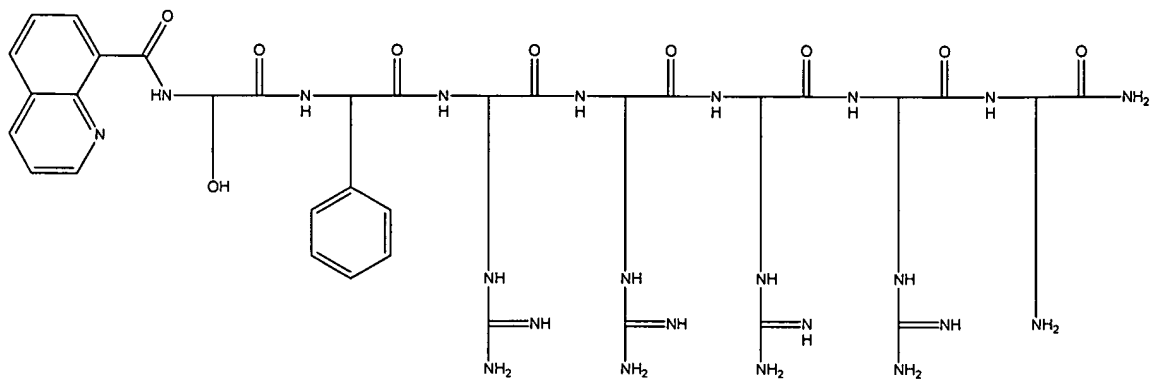
397236.1

Applicant: David S. Lawrence

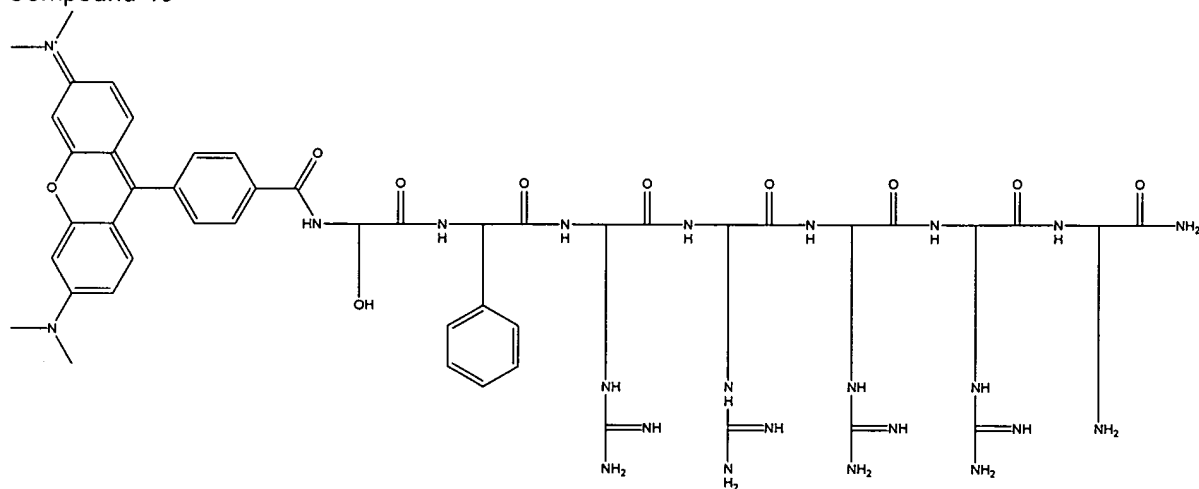
Serial No.: 10/755,086

Filed: January 9, 2004

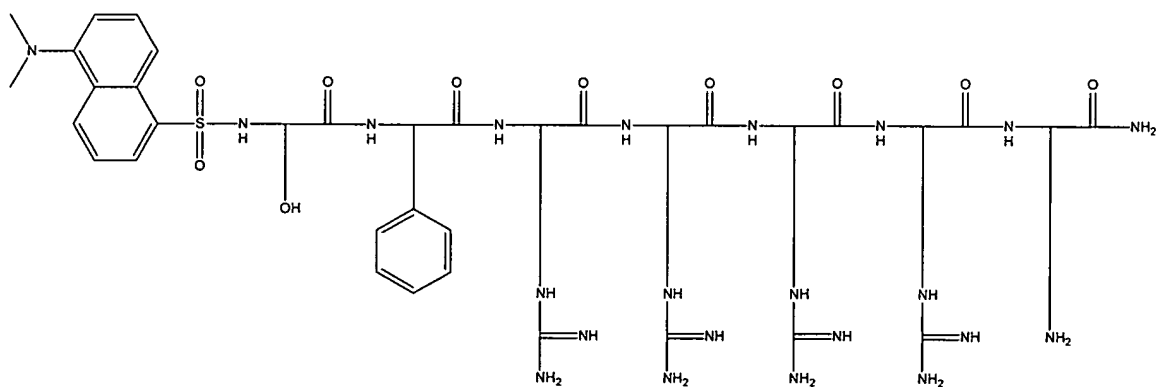
page 28 of 192



Compound 49

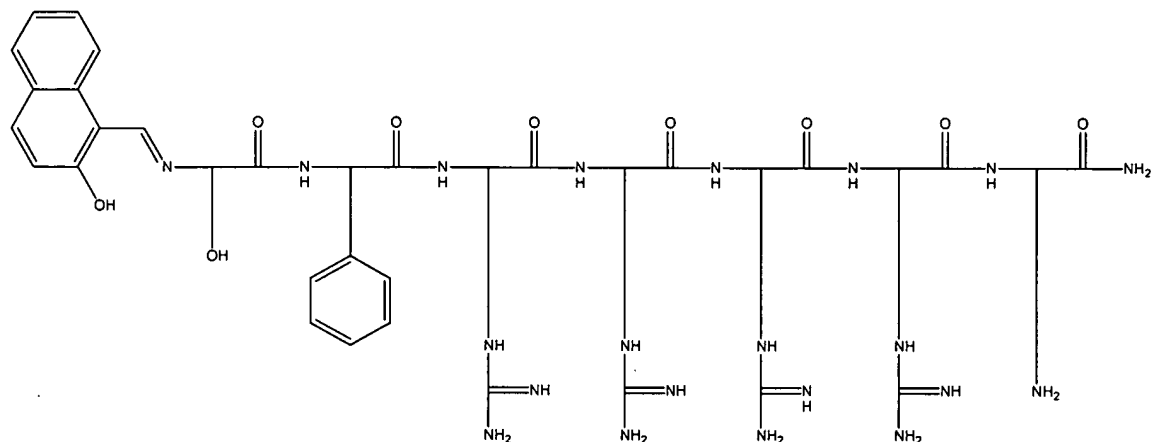


Compound 50

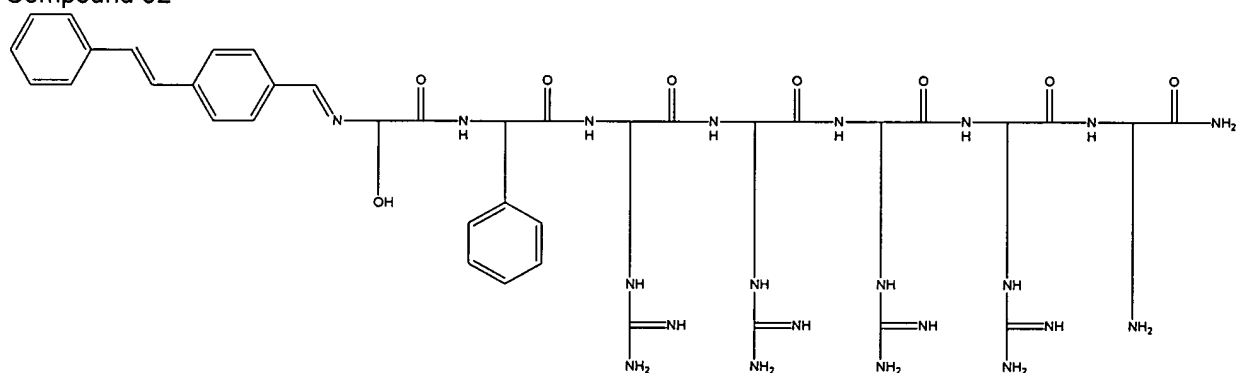


Compound 51

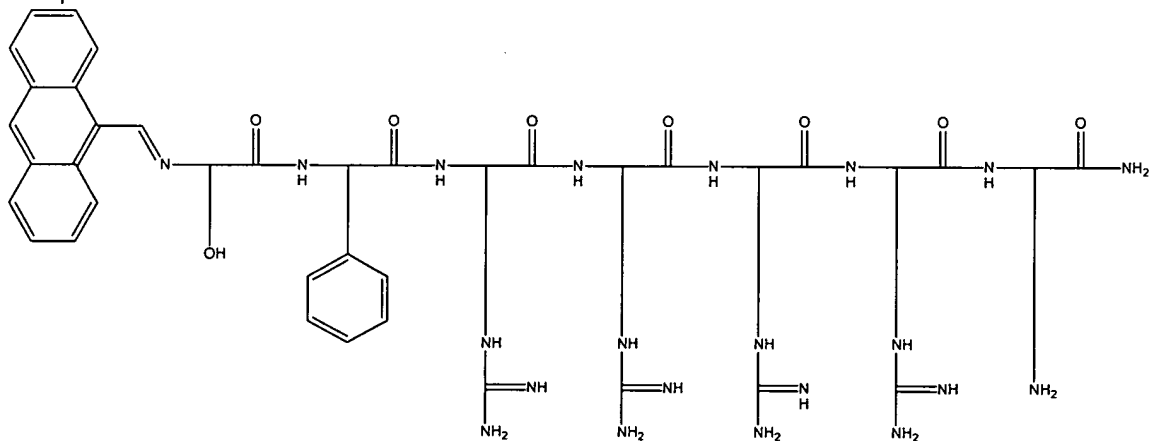
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 29 of 192



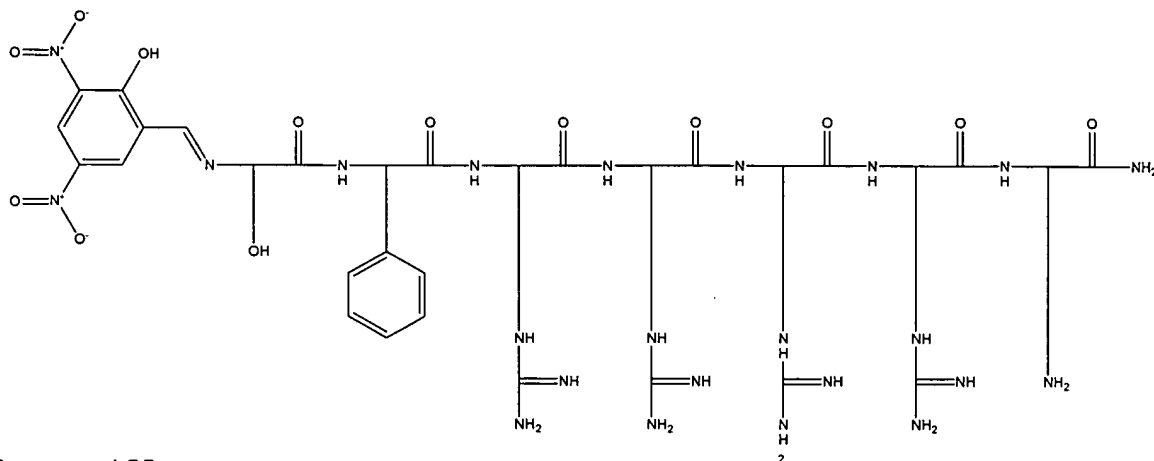
Compound 52



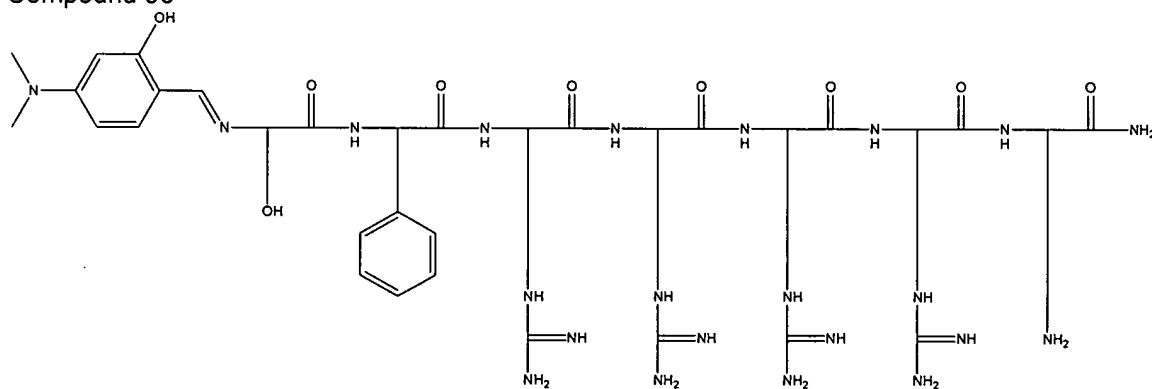
Compound 53



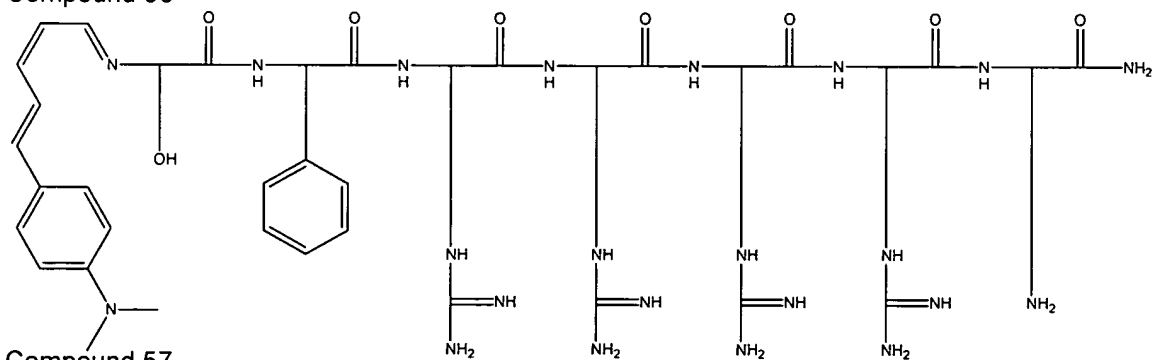
Compound 54



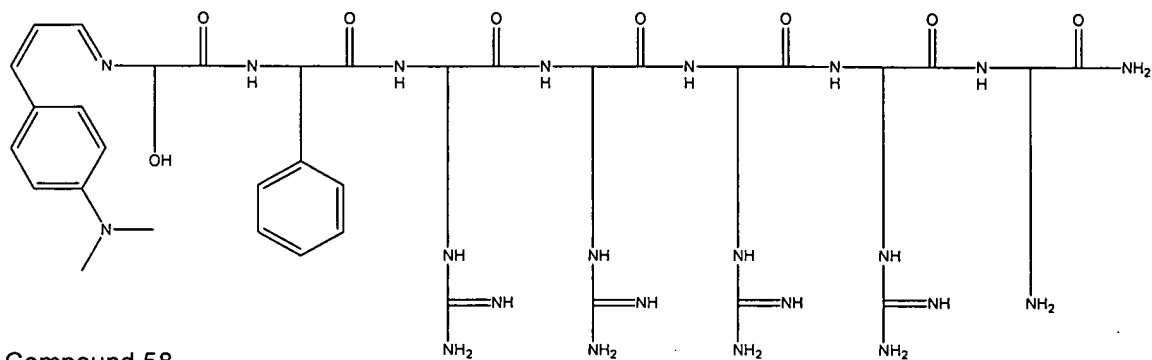
Compound 55



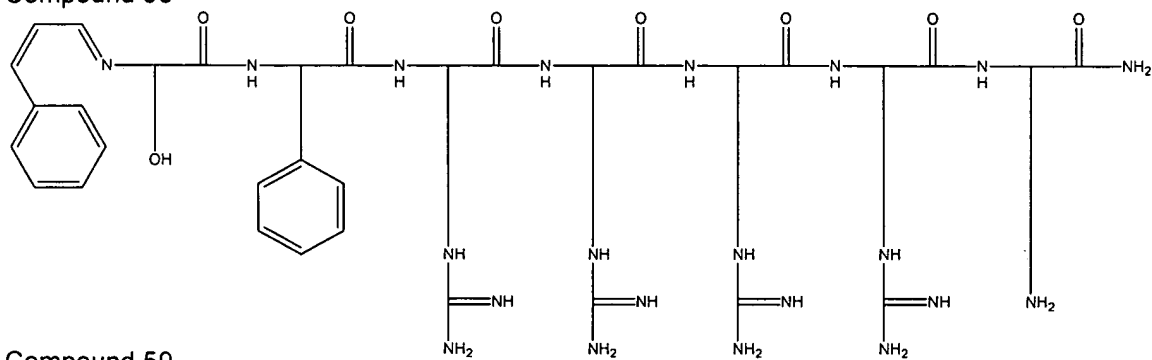
Compound 56



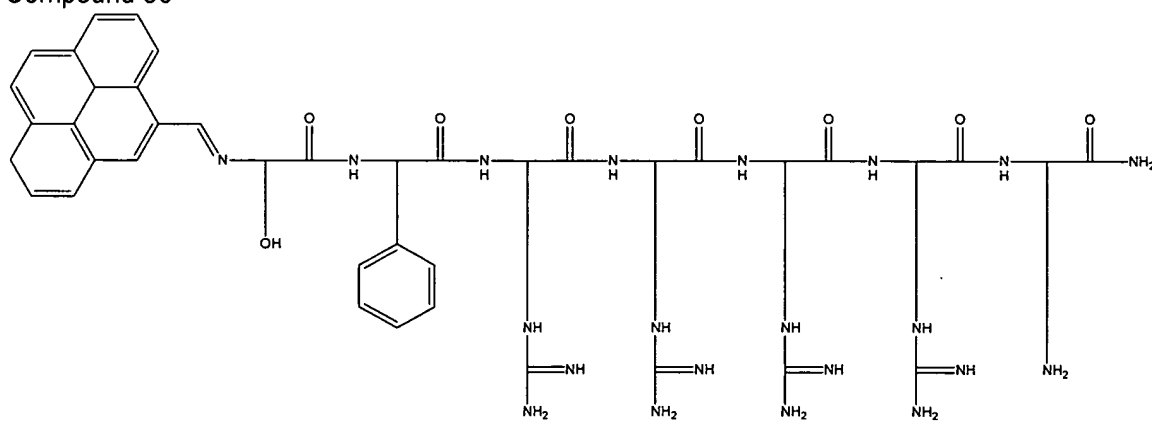
Compound 57



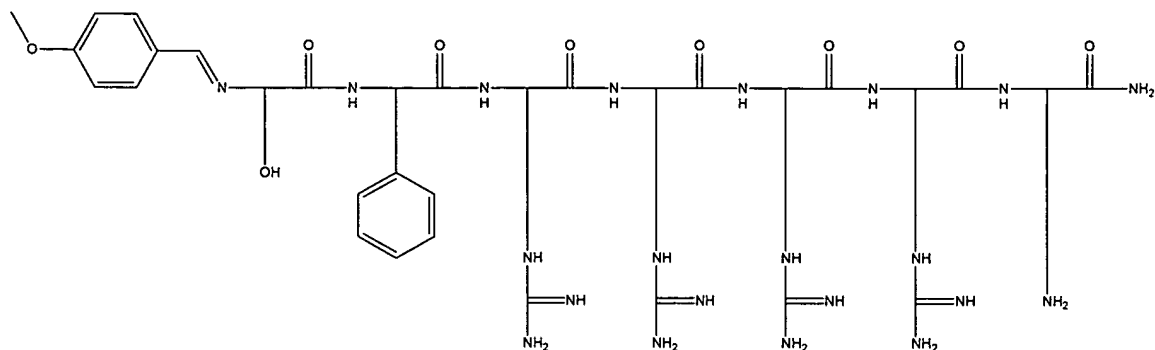
Compound 58



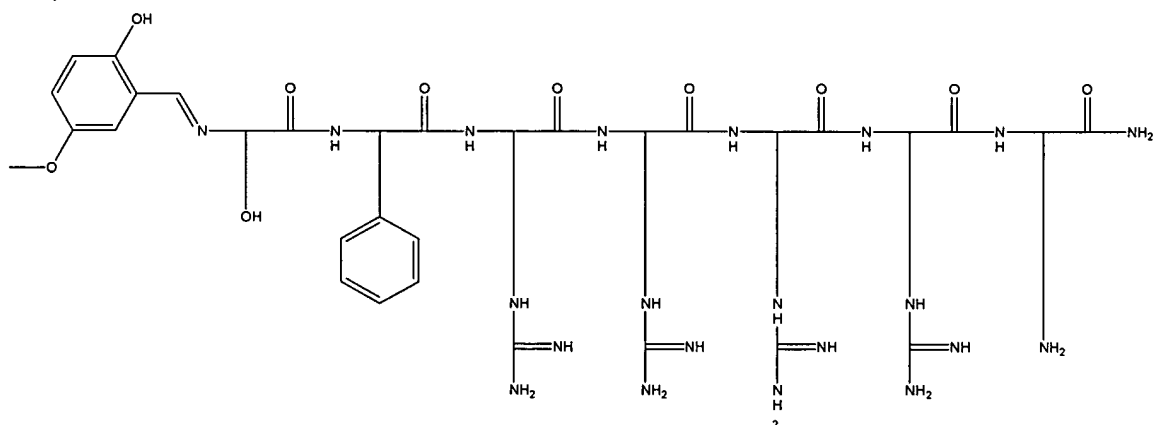
Compound 59



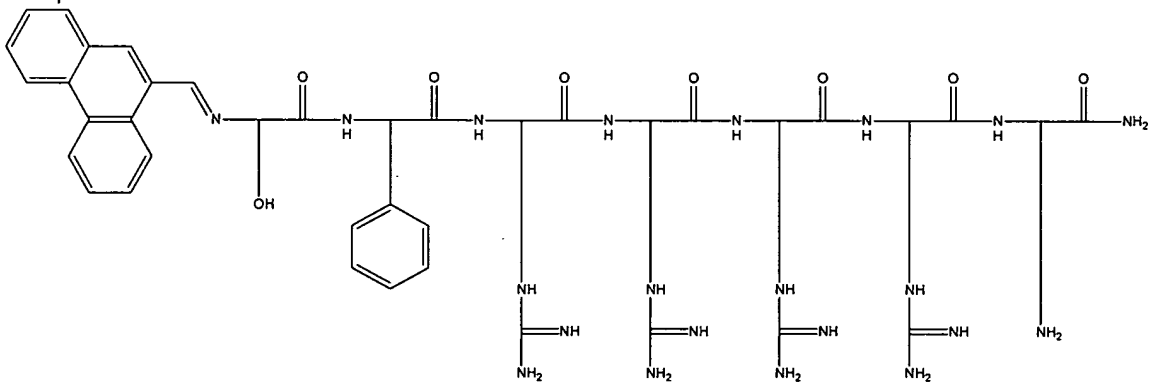
Compound 60



Compound 61

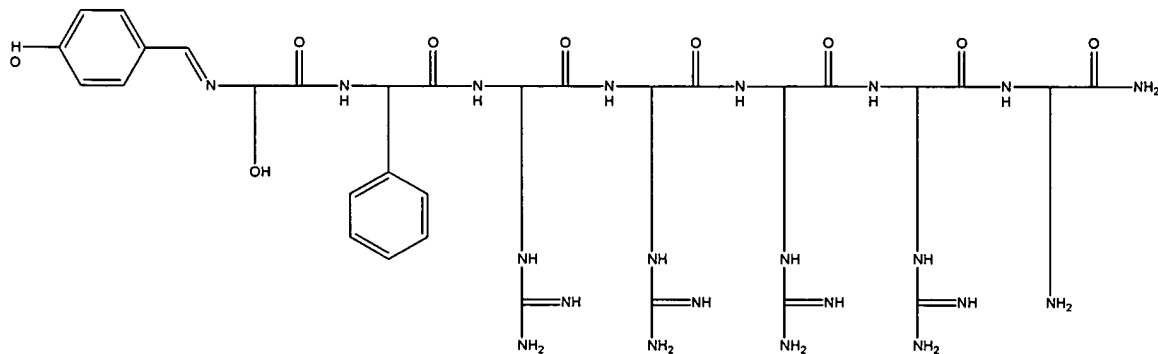


Compound 62

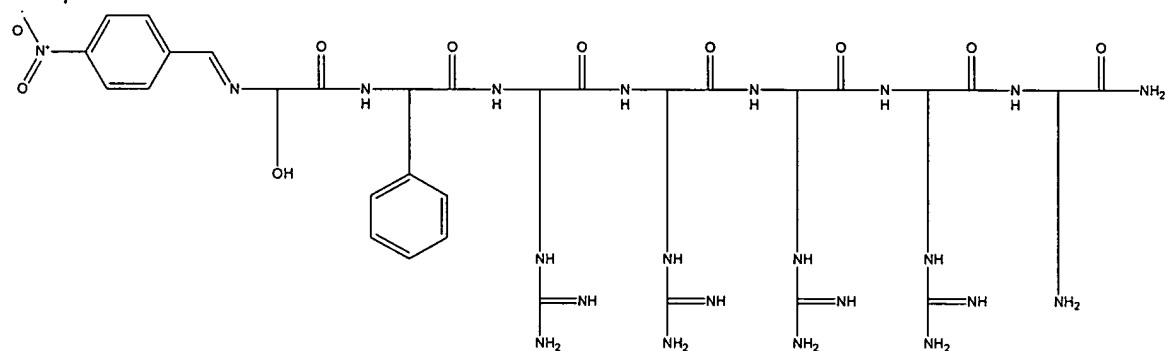


Compound 63

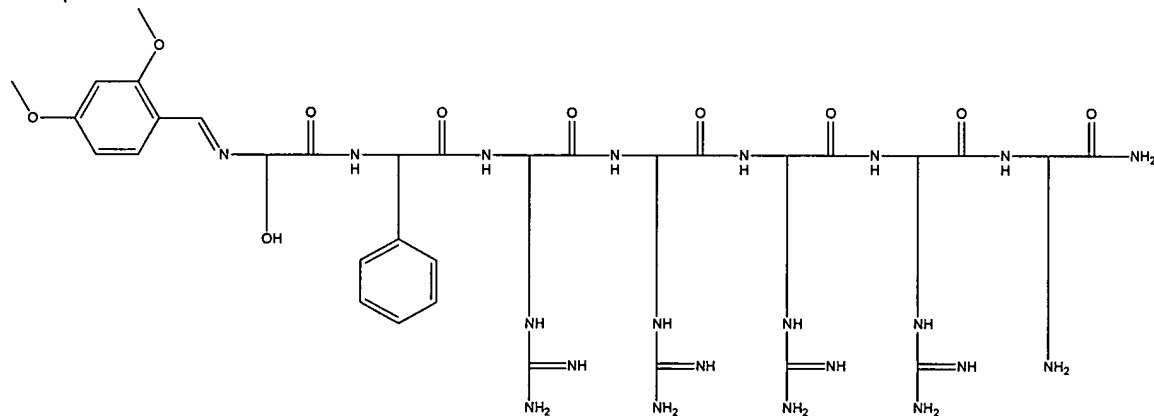
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 33 of 192



Compound 64

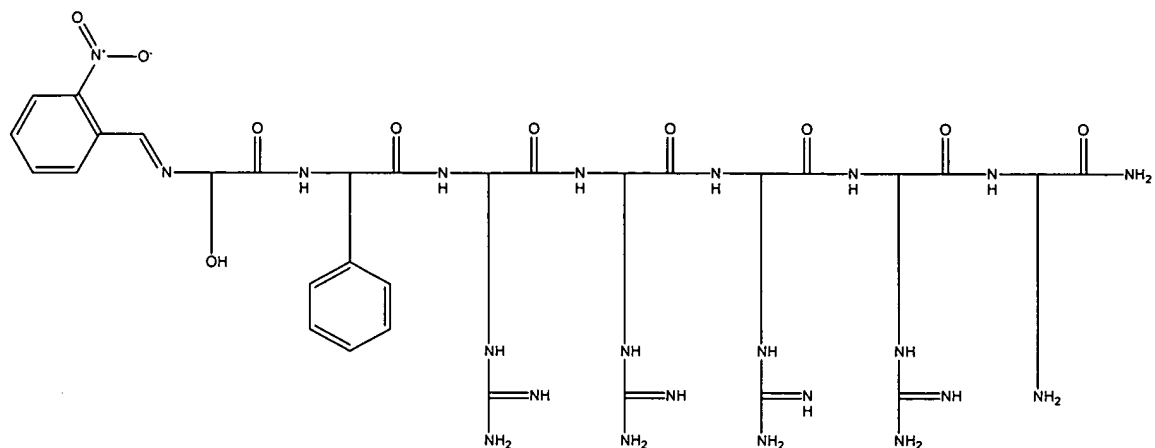


Compound 65

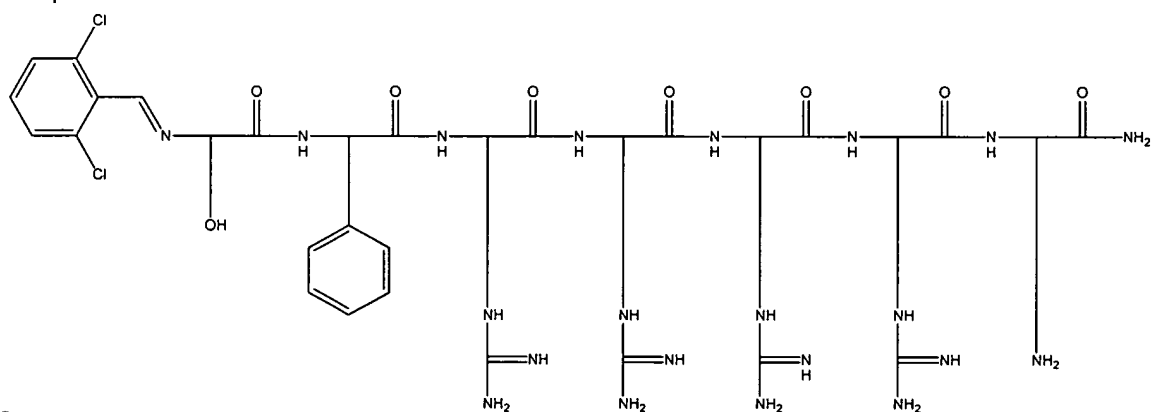


Compound 66

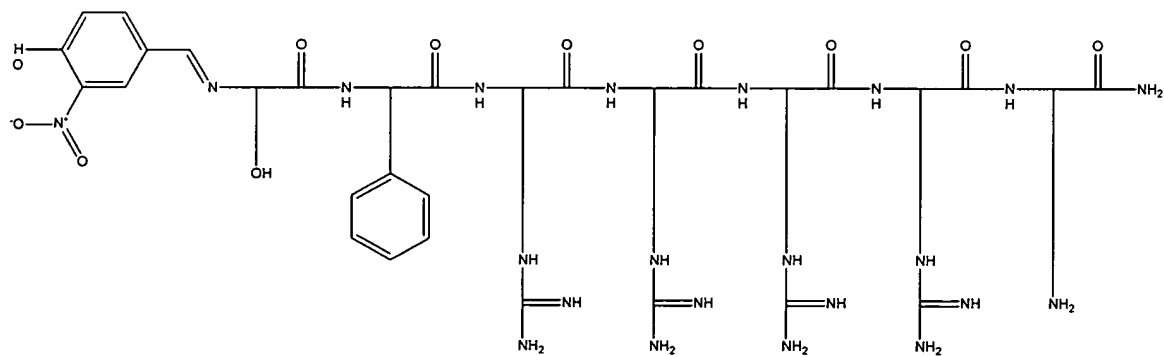
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 34 of 192



Compound 67

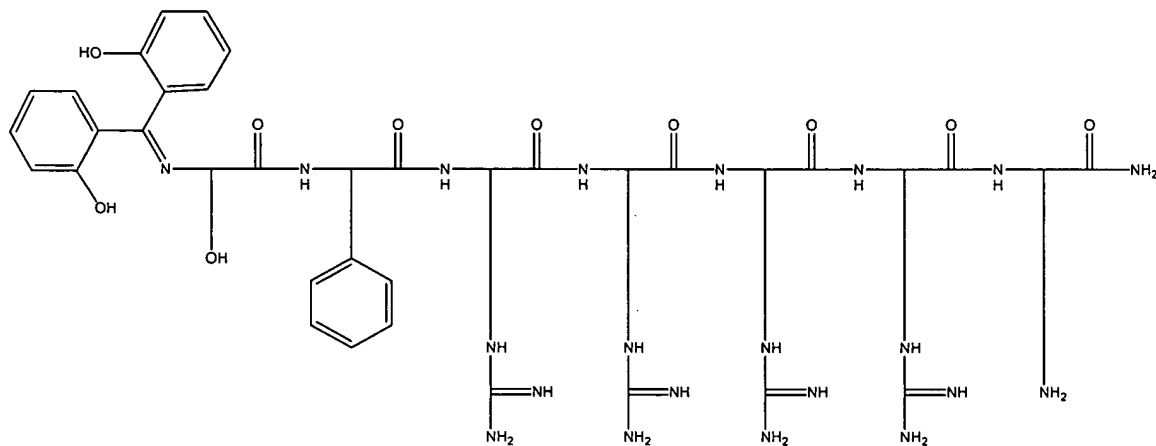


Compound 68

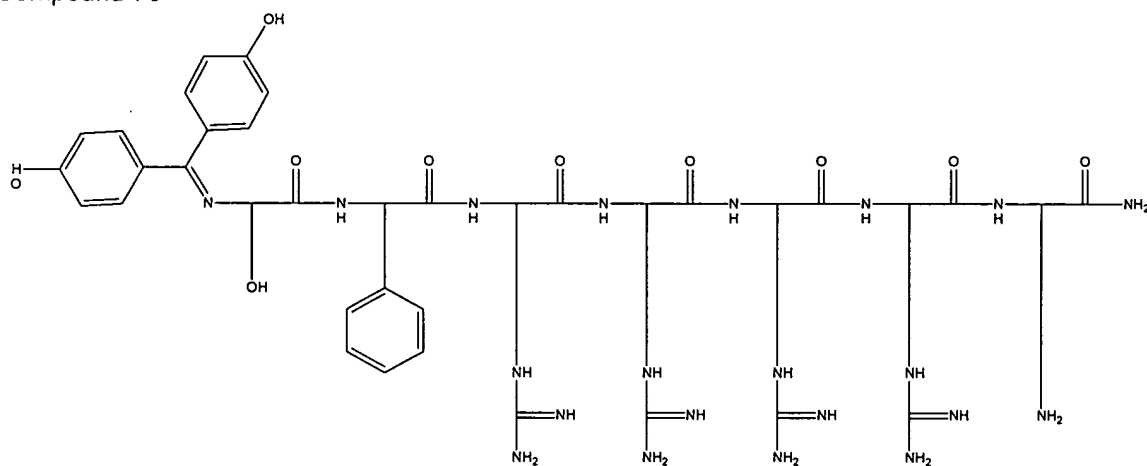


Compound 69

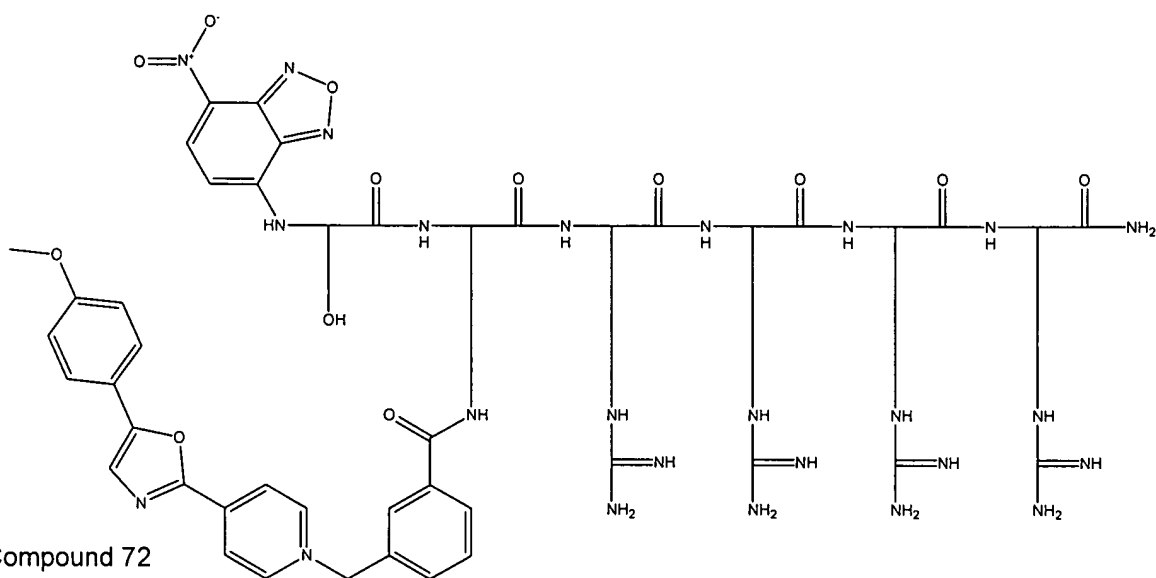
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 35 of 192



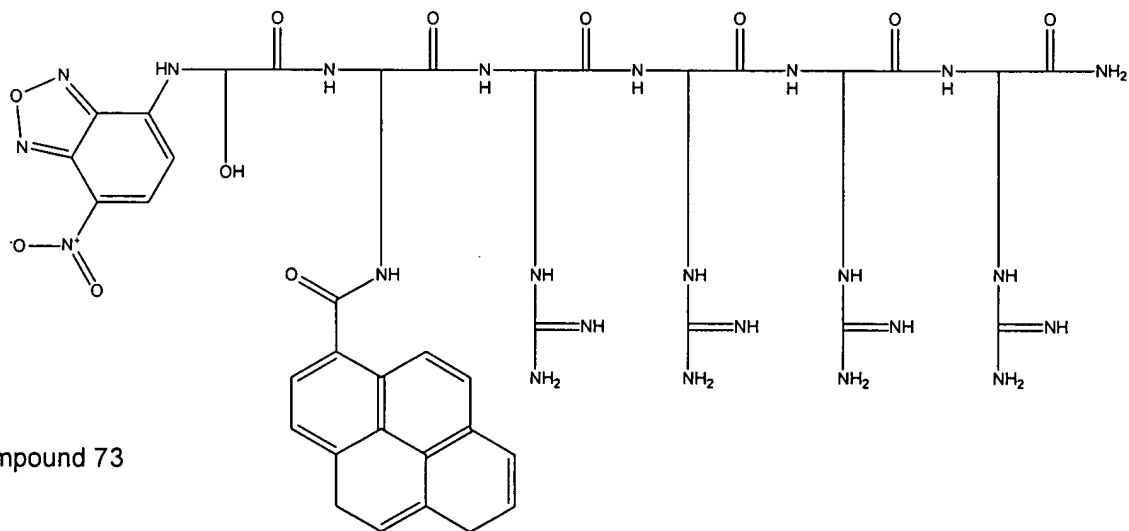
Compound 70



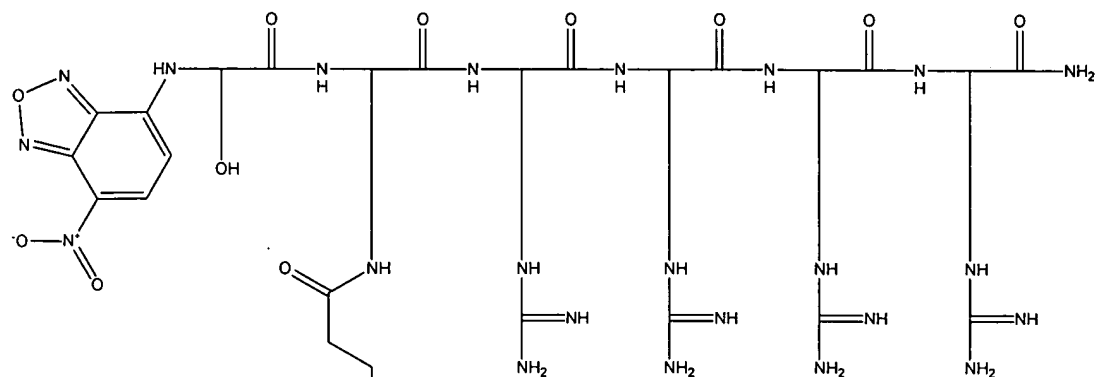
Compound 71



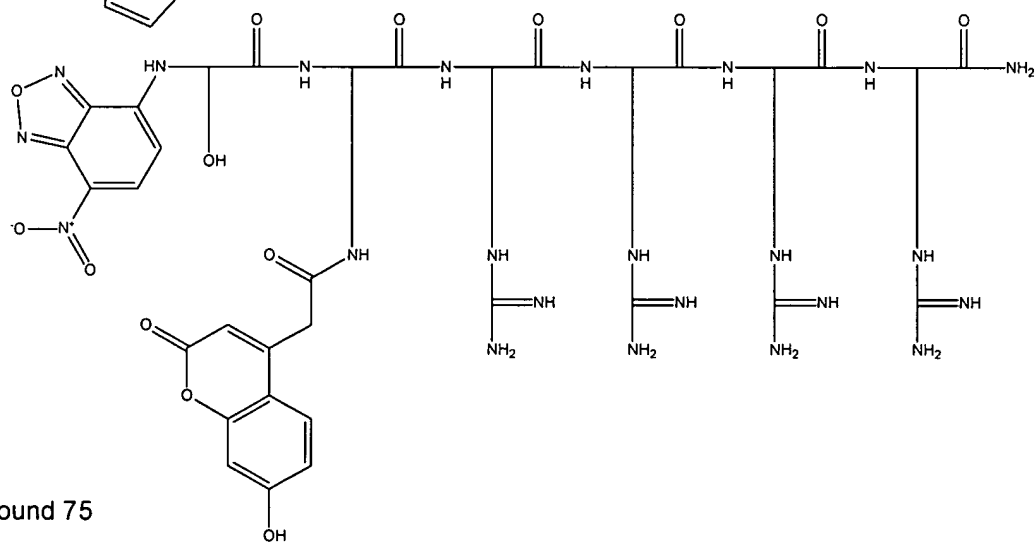
Compound 72



Compound 73

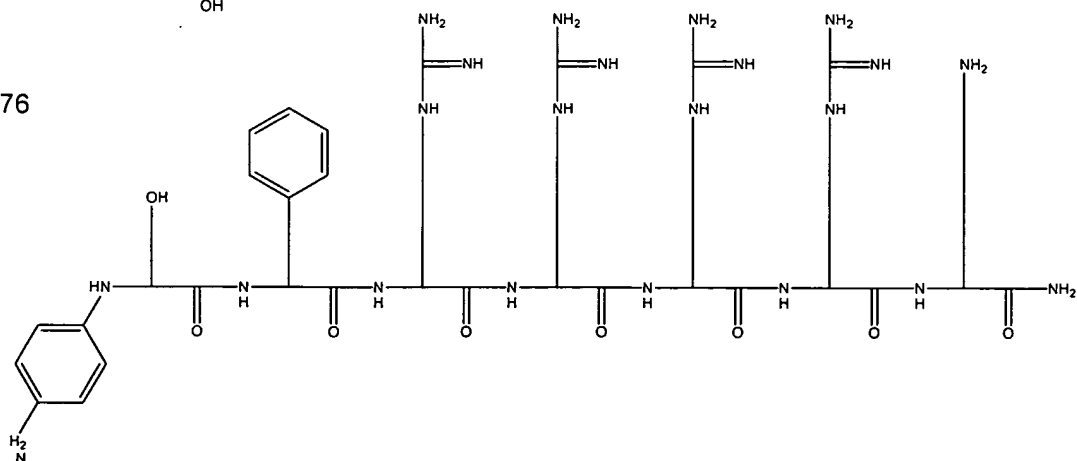


Compound 74

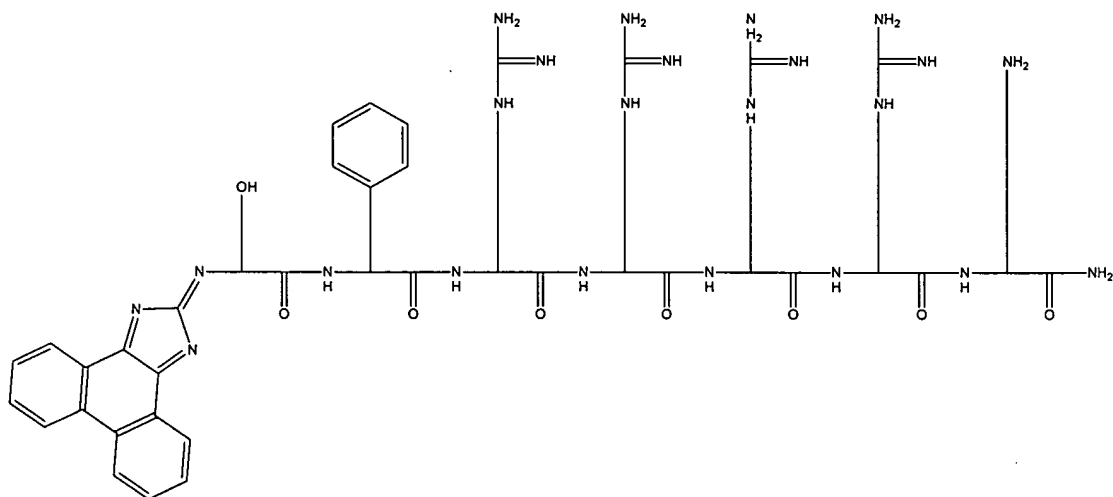


Compound 75

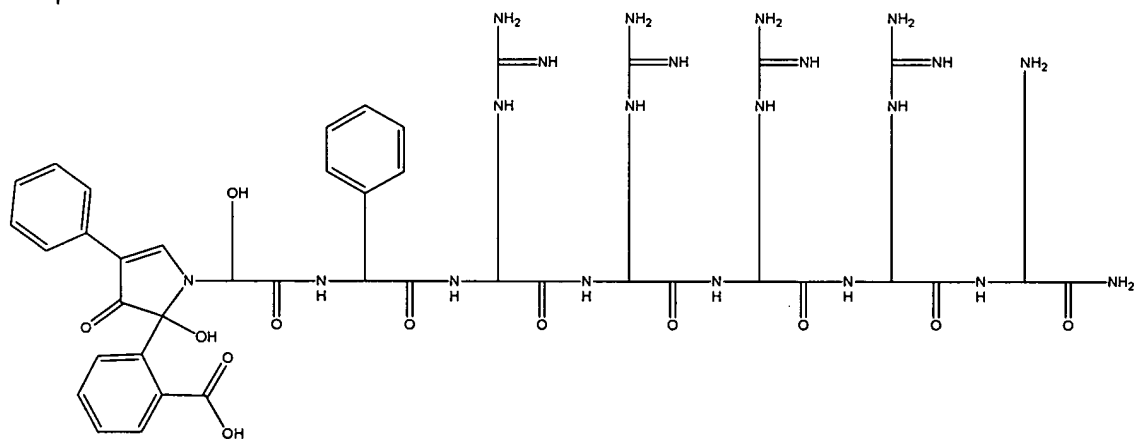
Compound 76



Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 38 of 192



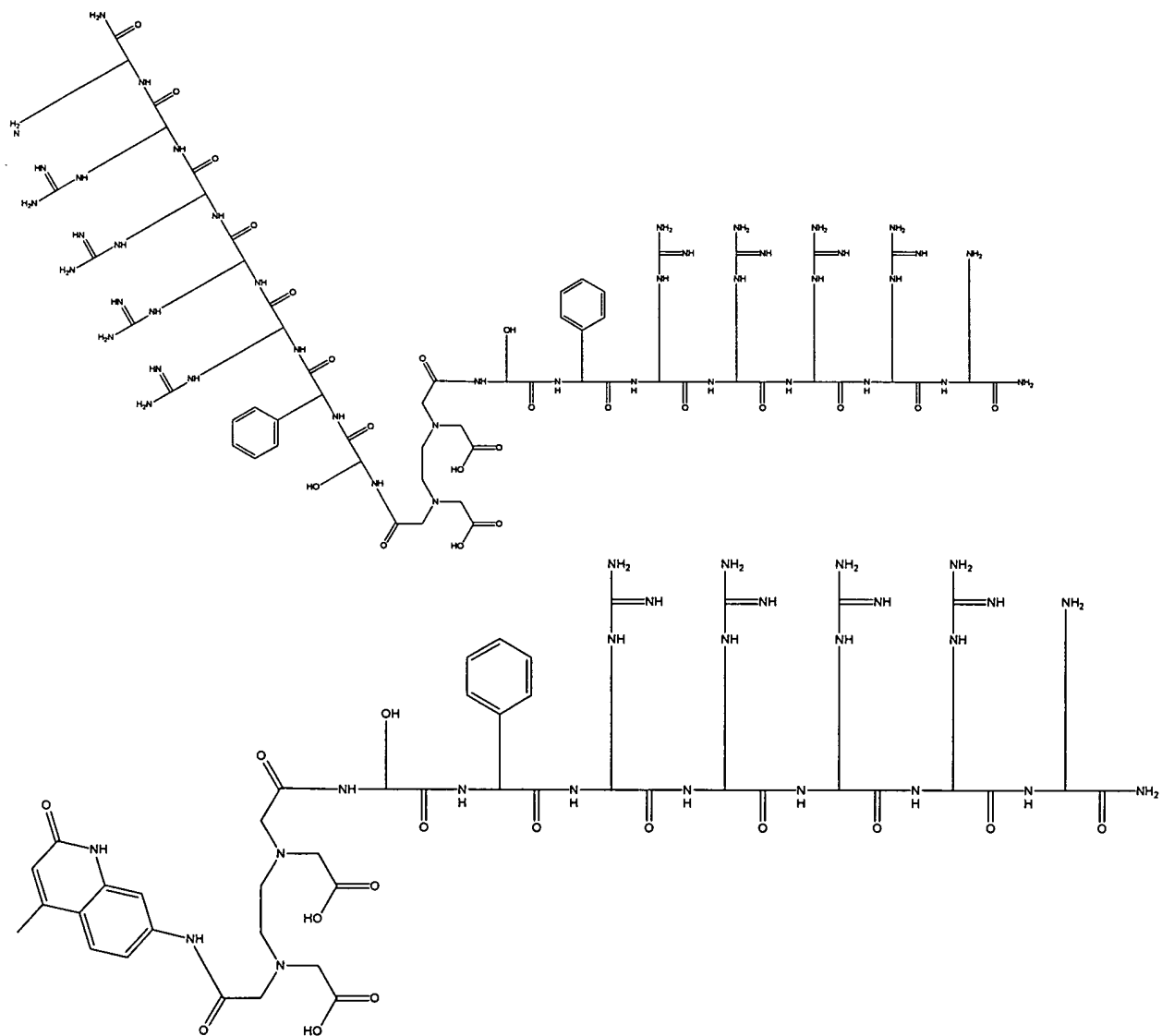
Compound 77



Compound 78

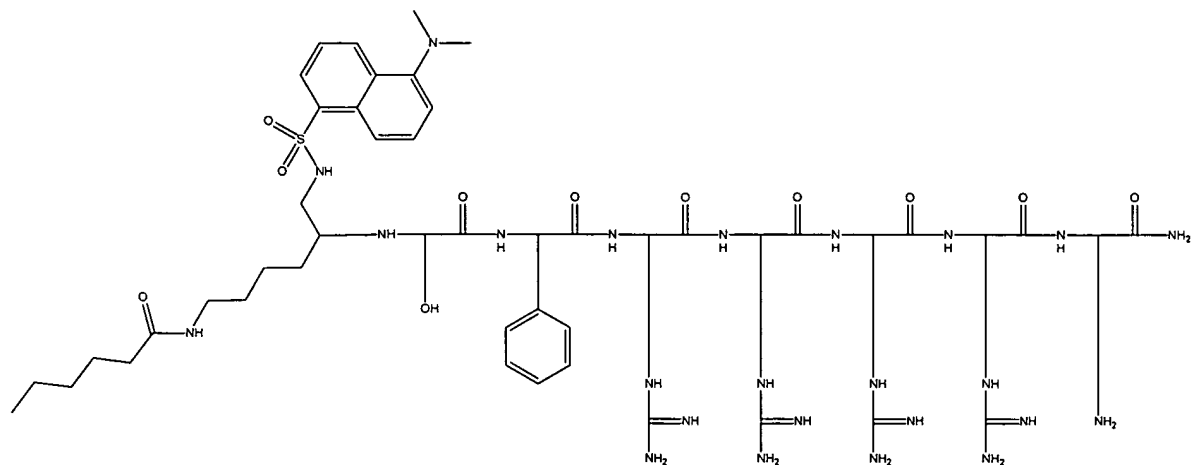
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 39 of 192

Compound 79



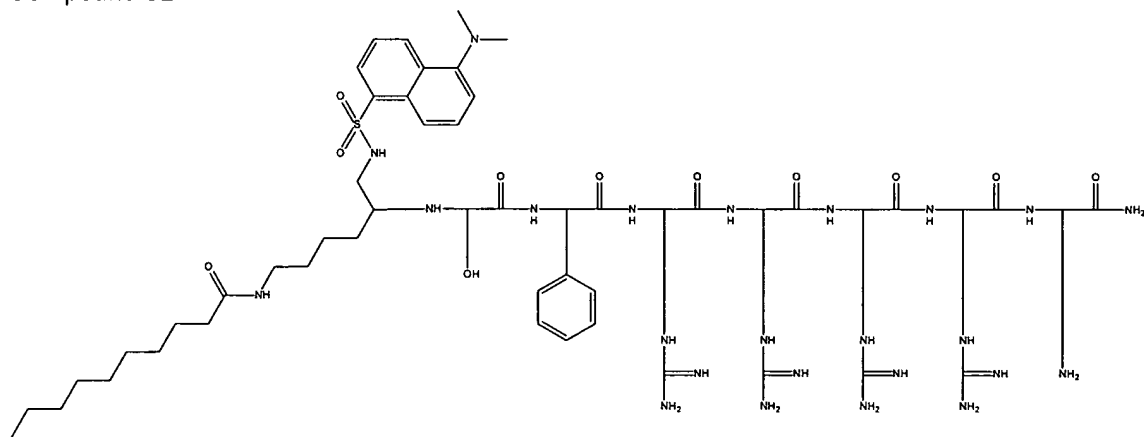
Compound 80

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 40 of 192

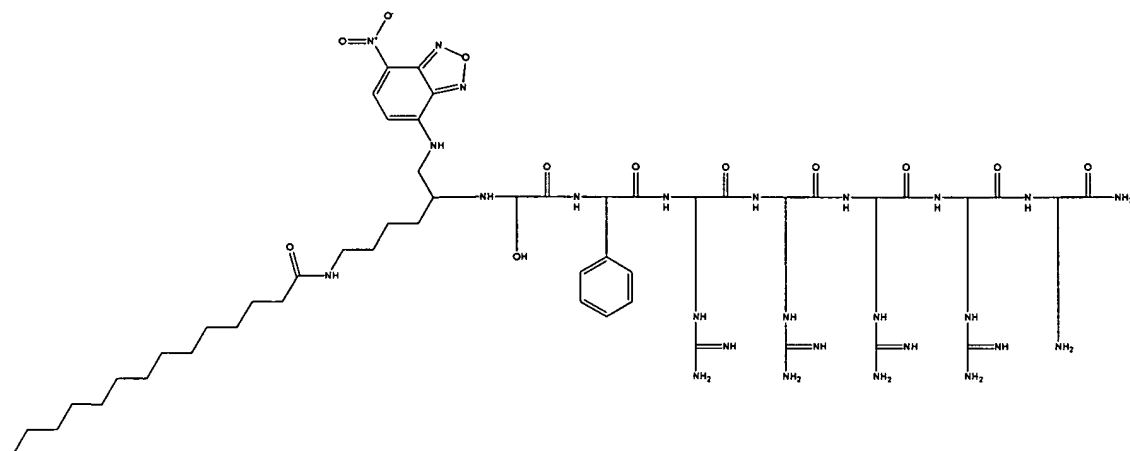


Compound 81

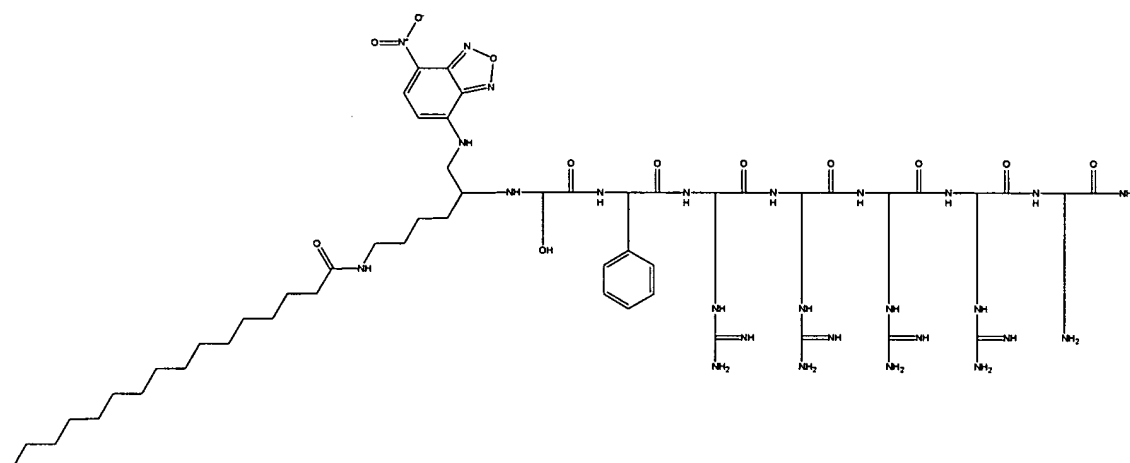
Compound 82



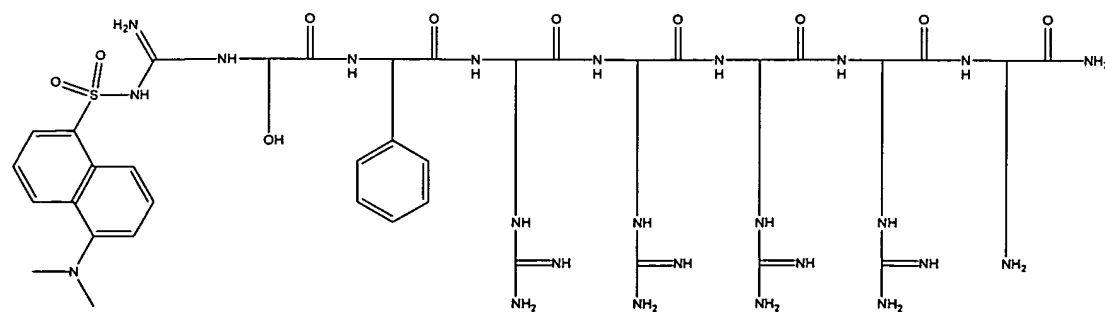
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 41 of 192



Compound 83

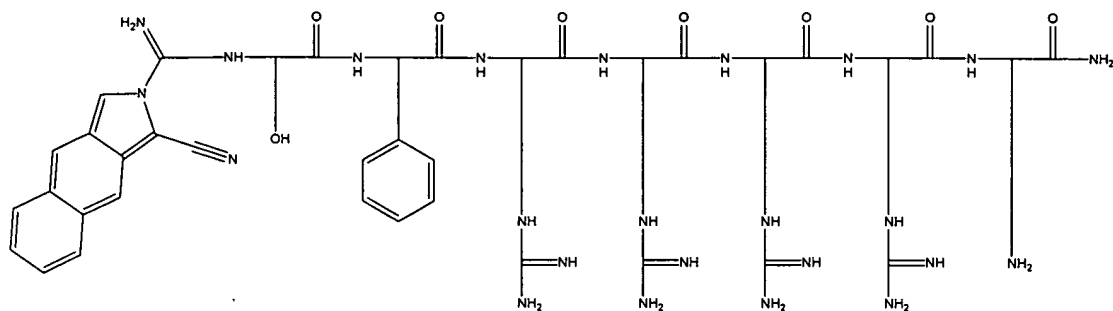


Compound 84

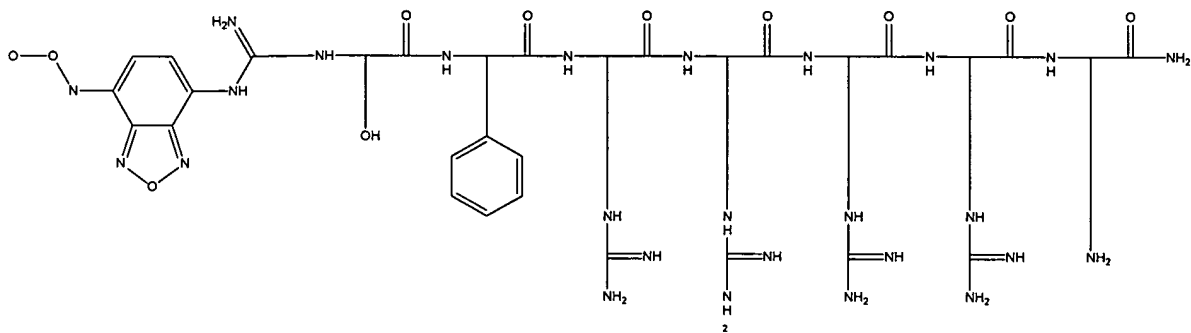


Compound 85

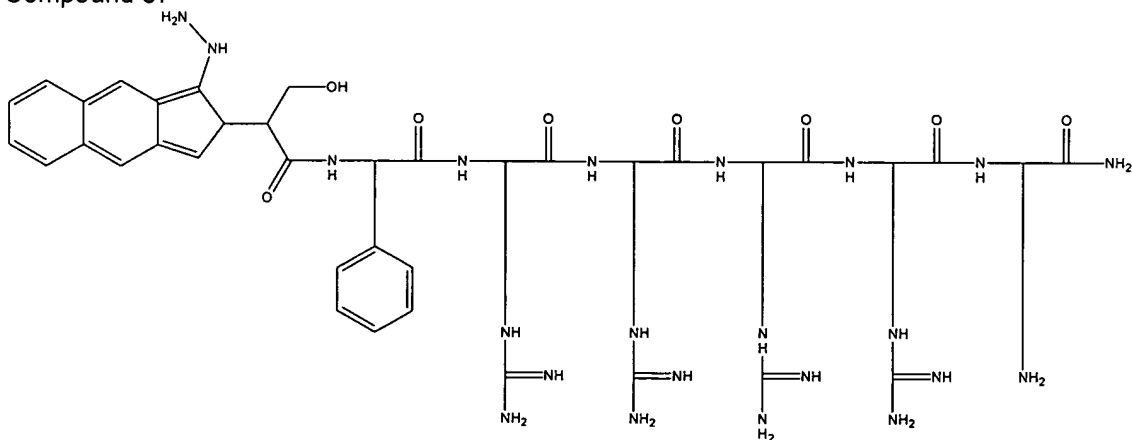
page 42 of 192



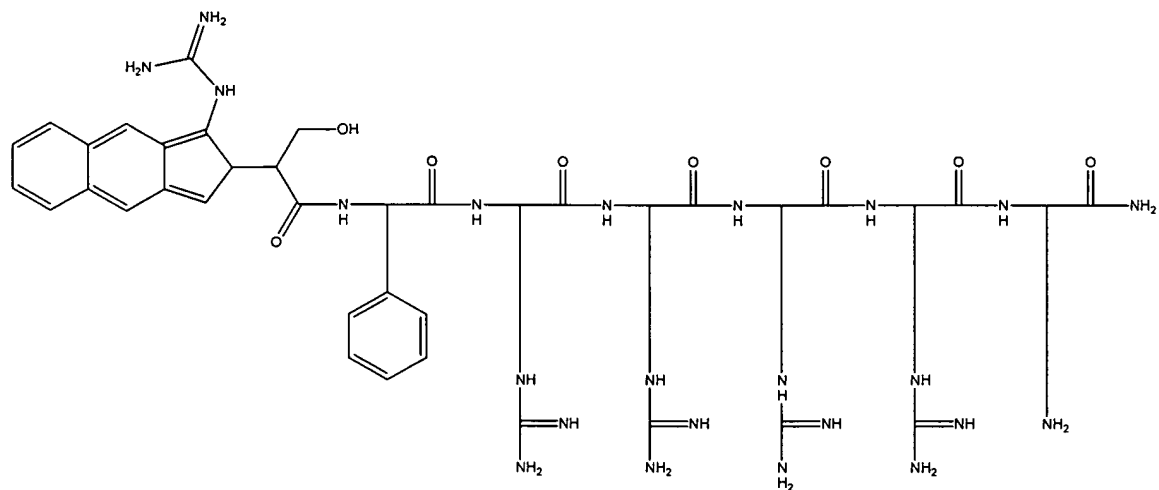
Compound 86



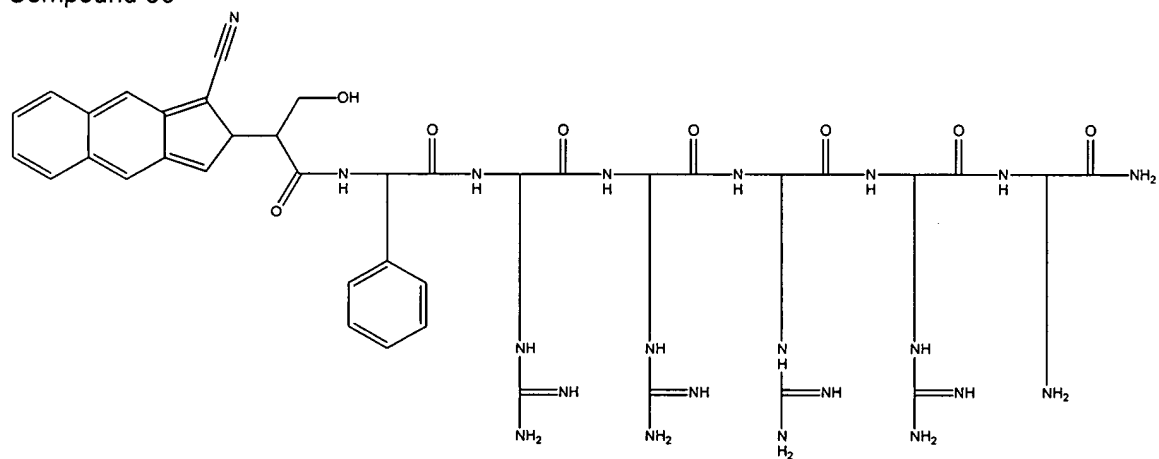
Compound 87



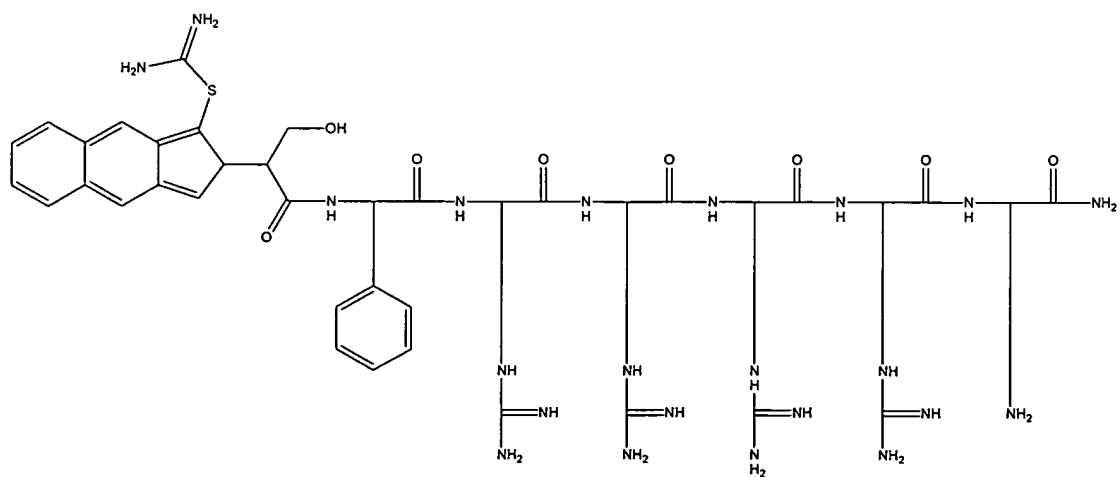
Compound 88



Compound 89

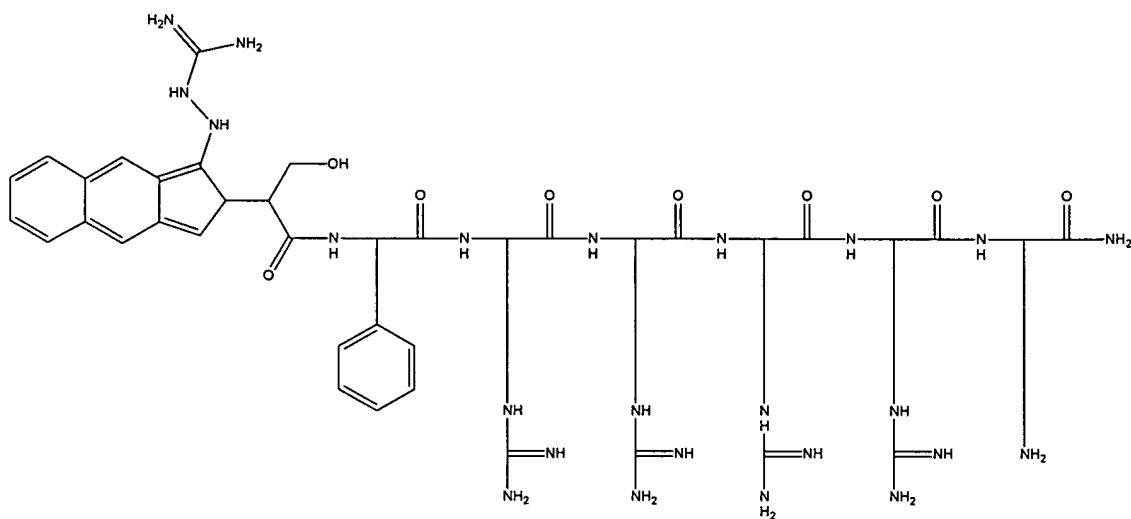


Compound 90

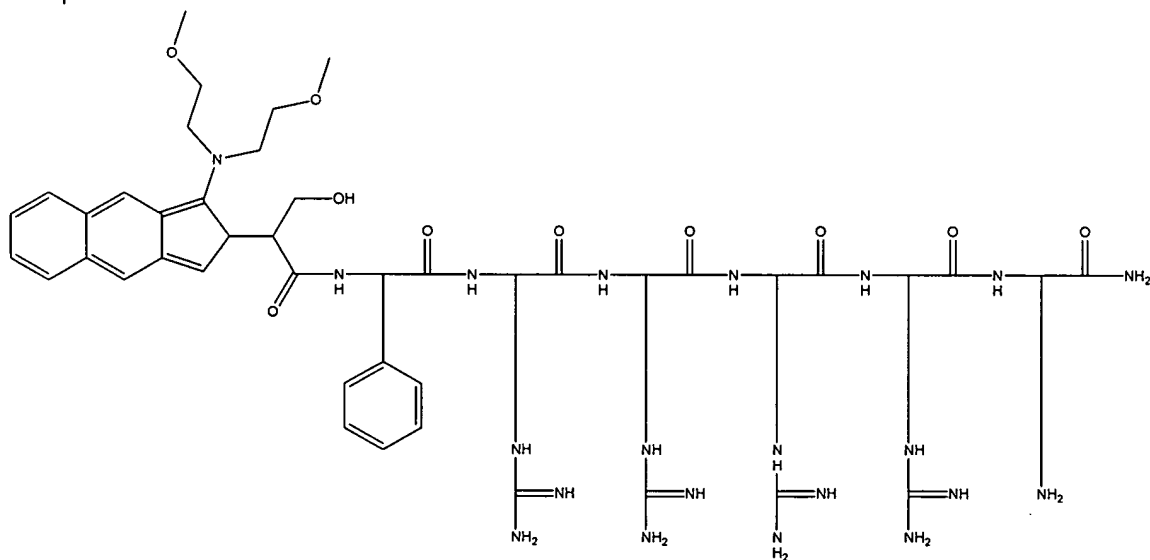


Compound 91

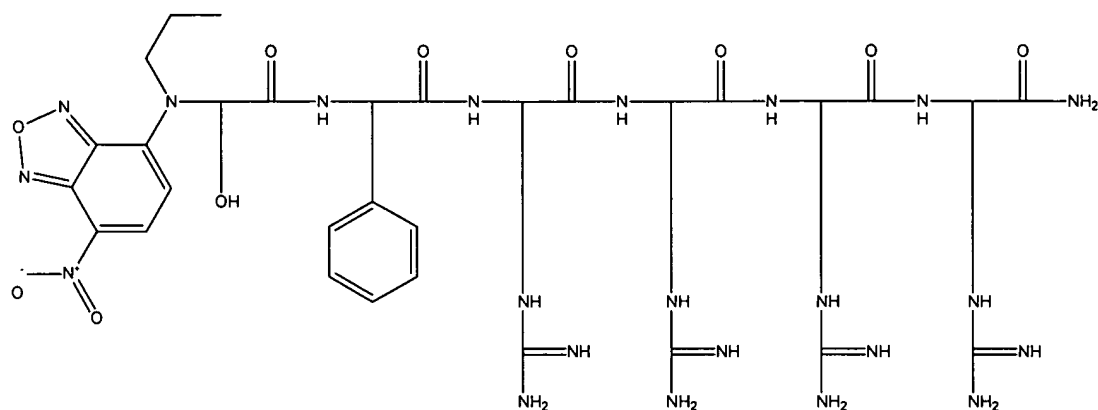
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 44 of 192



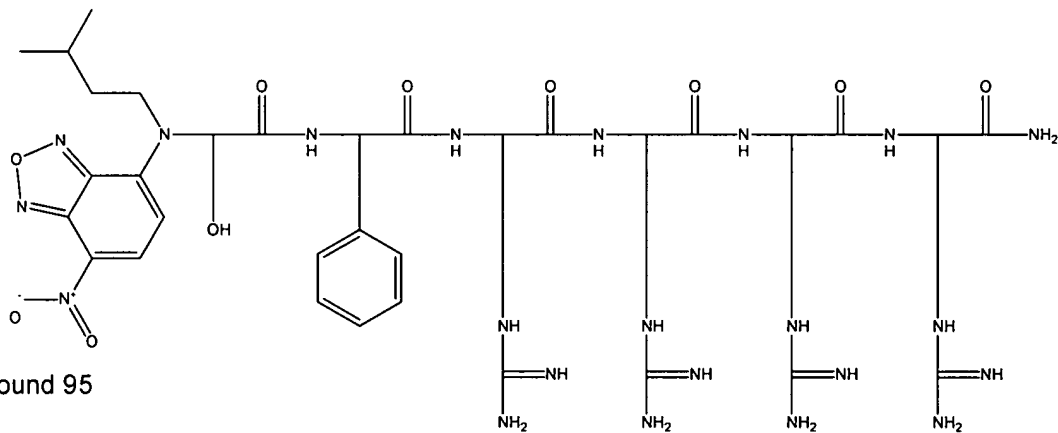
Compound 92



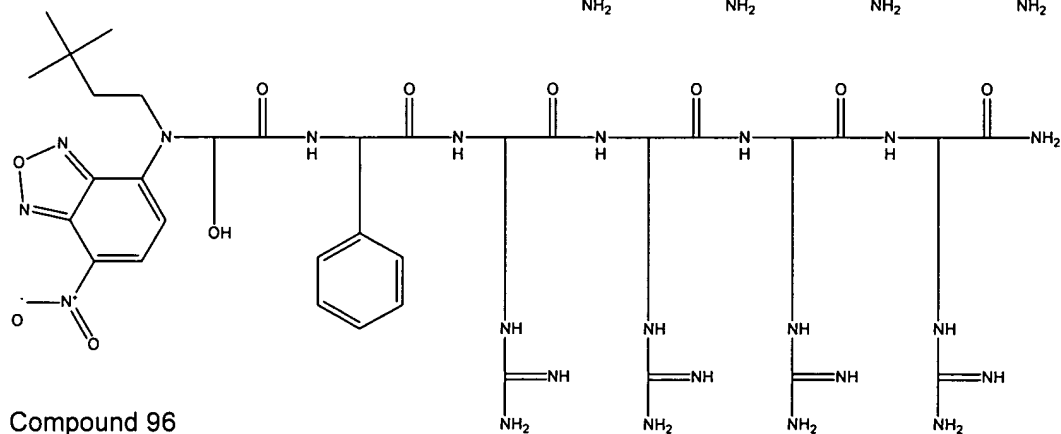
Compound 93



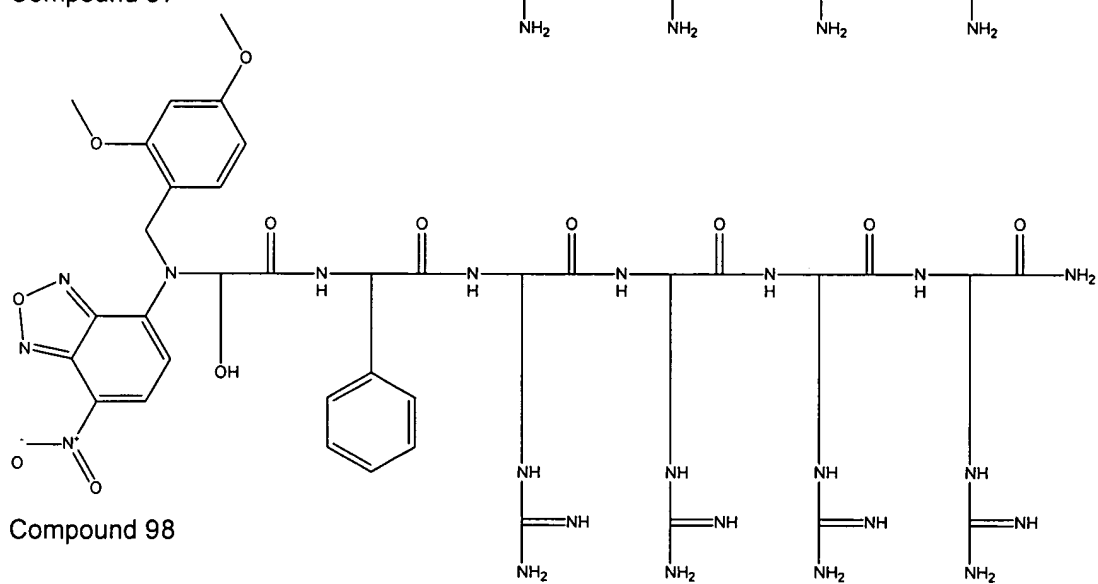
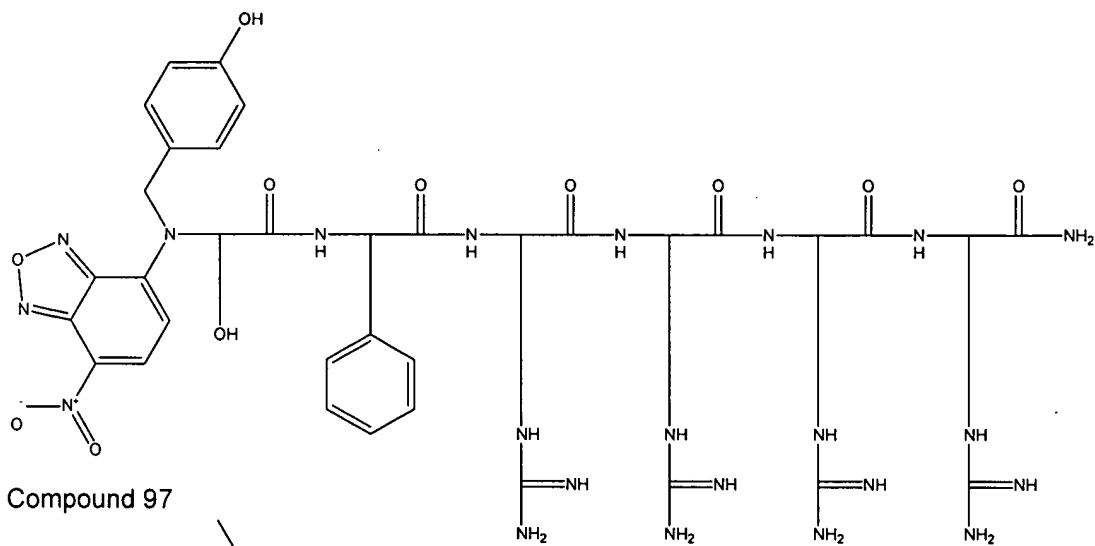
Compound 94

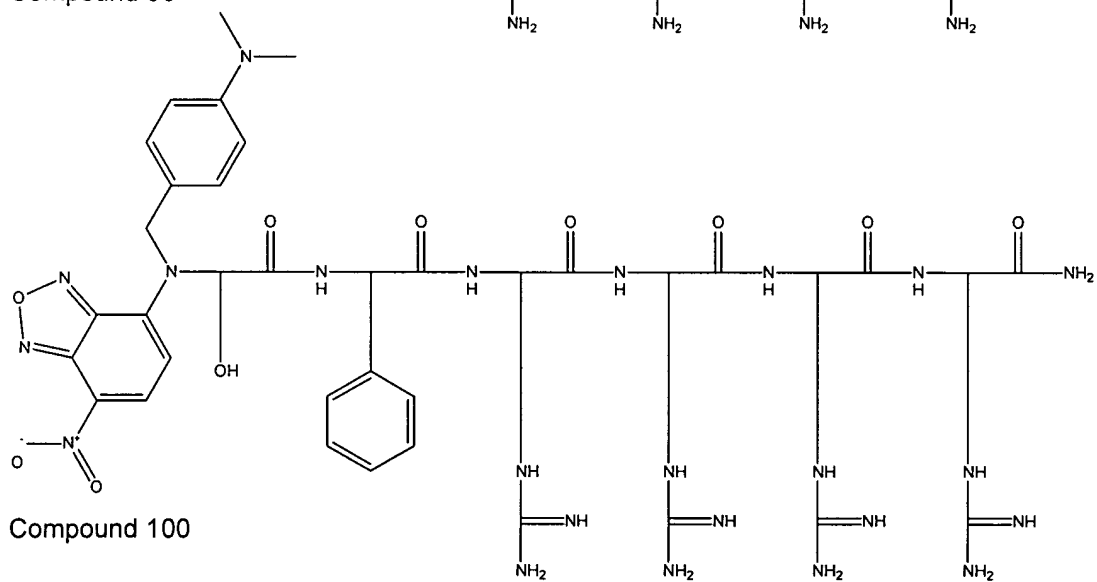
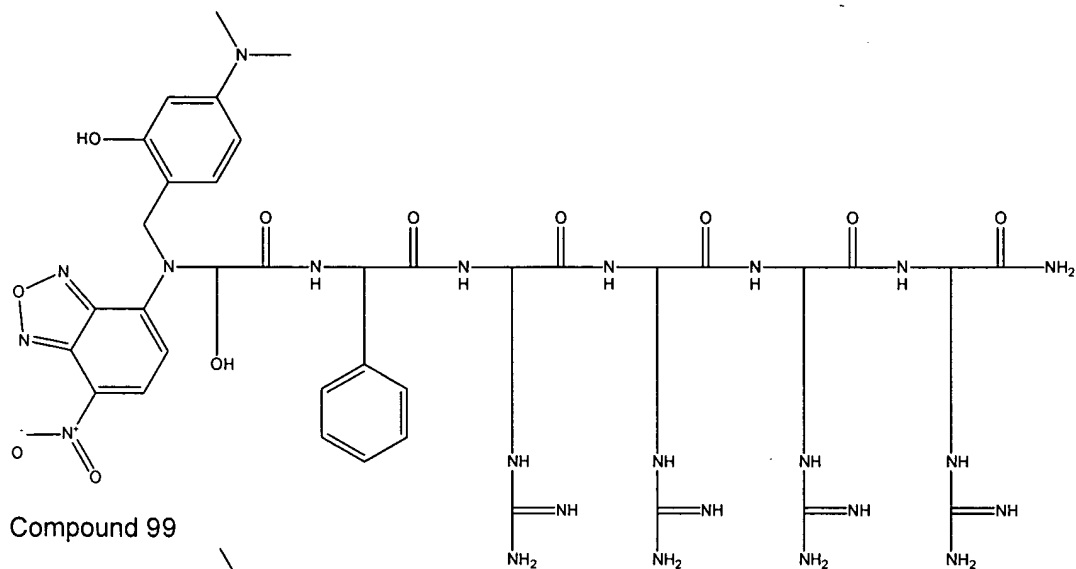


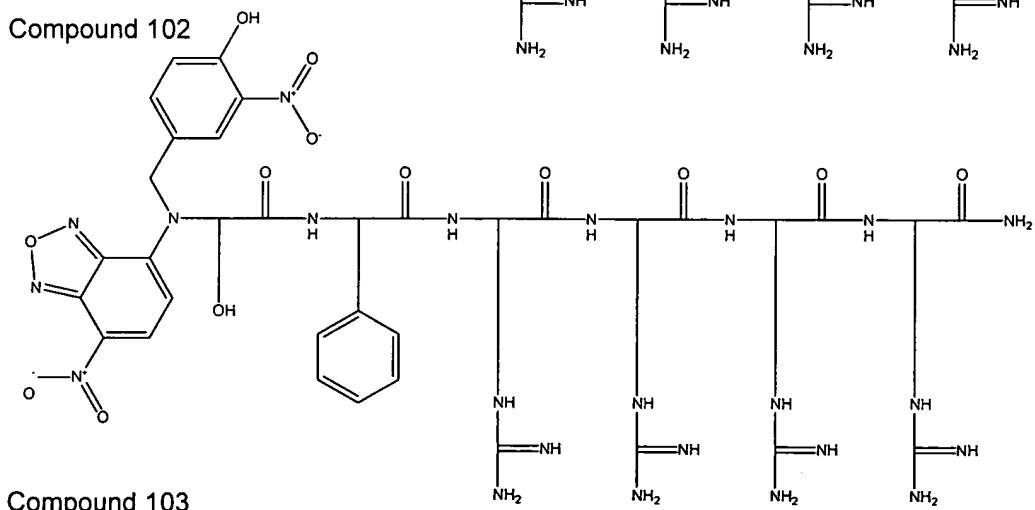
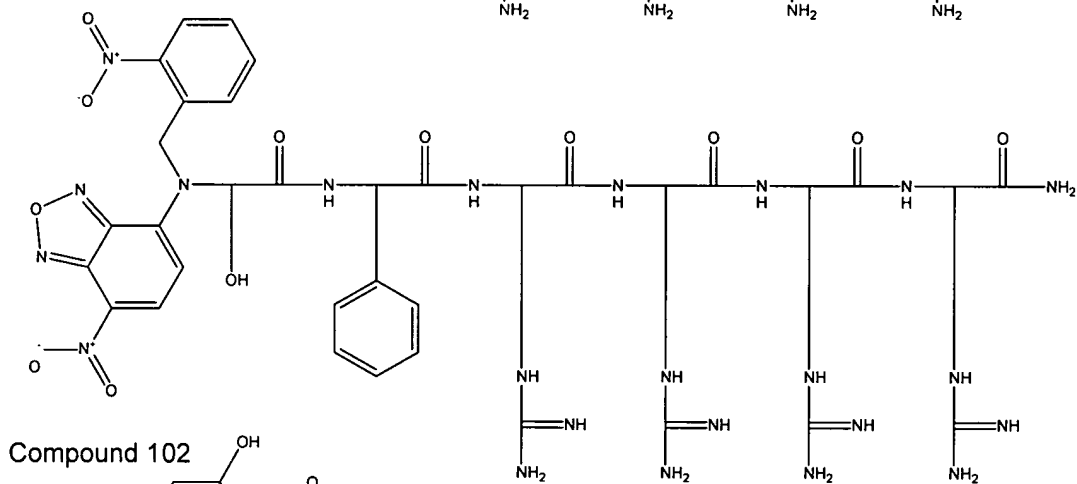
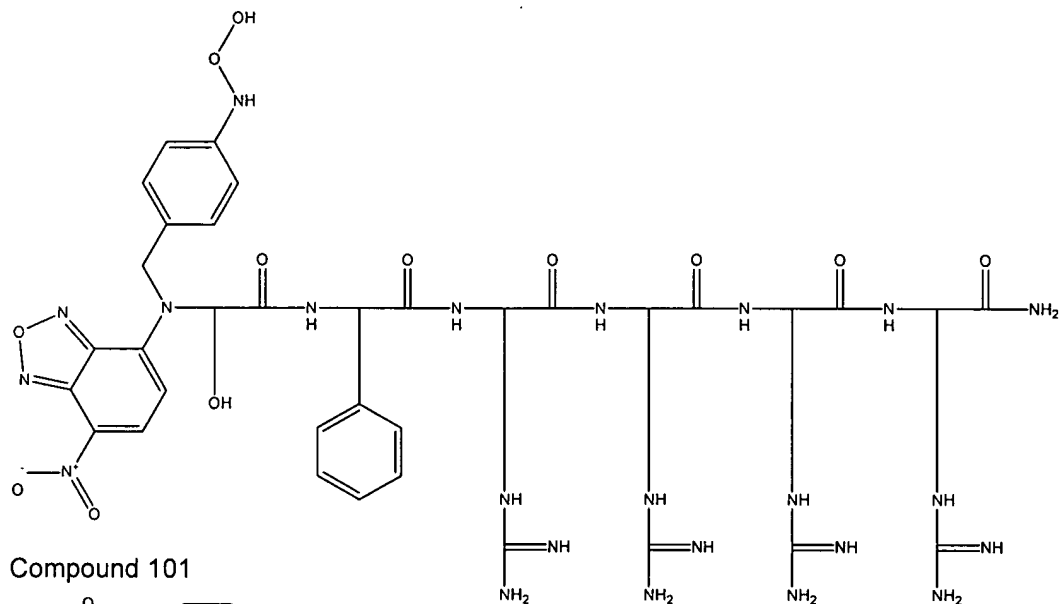
Compound 95



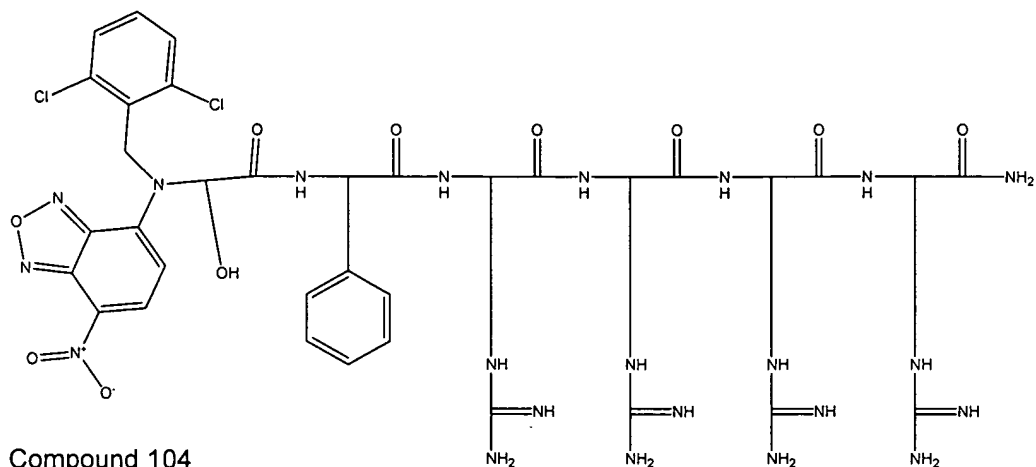
Compound 96



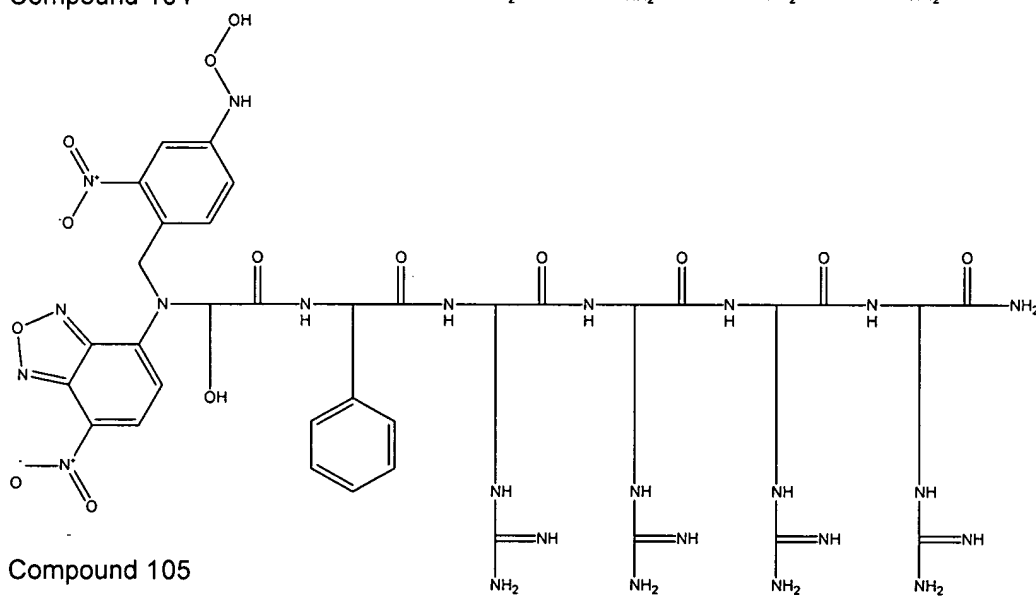




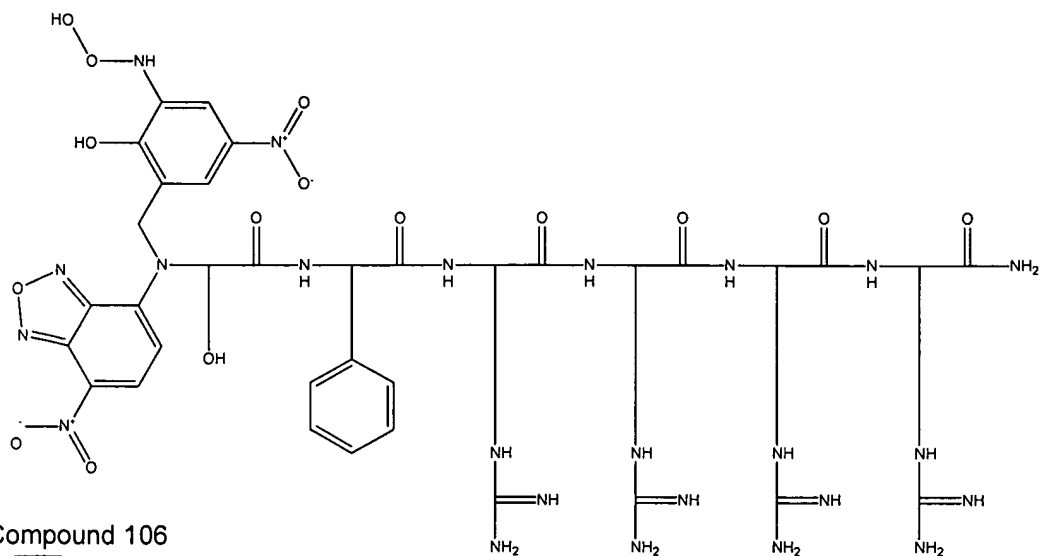
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 49 of 192



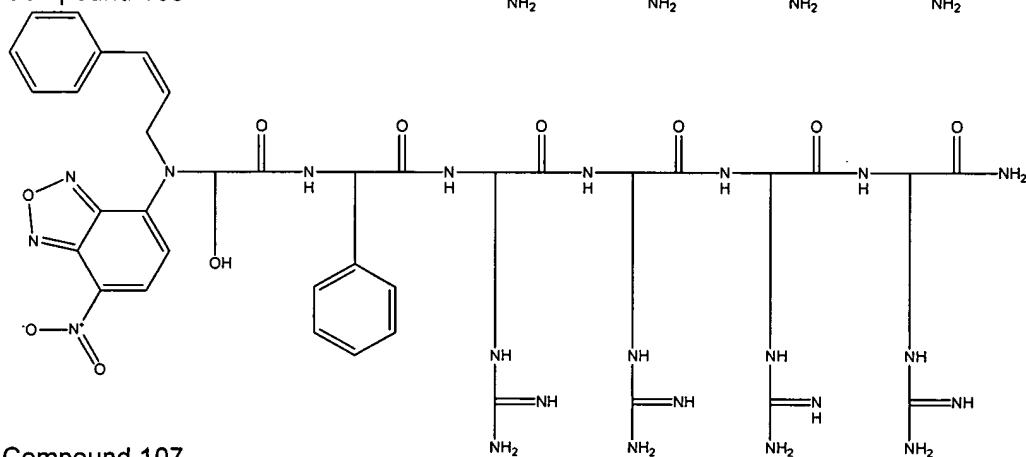
Compound 104



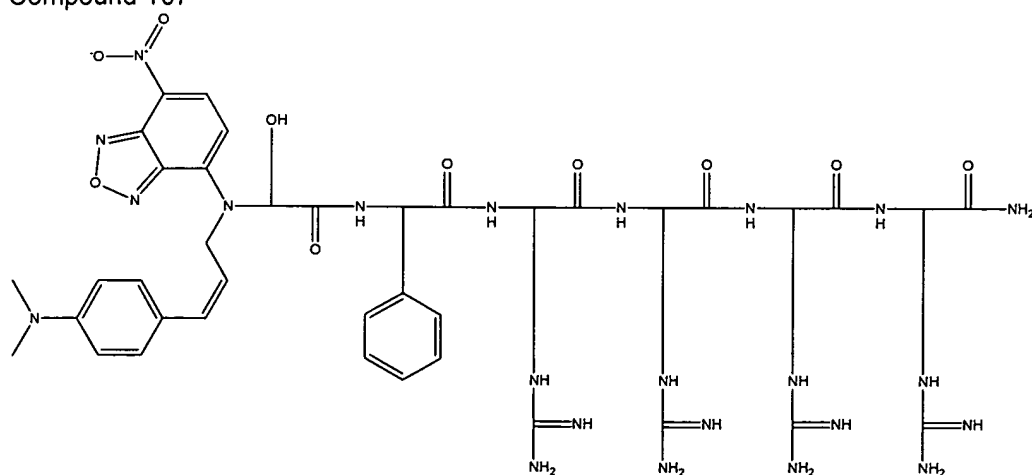
Compound 105



Compound 106

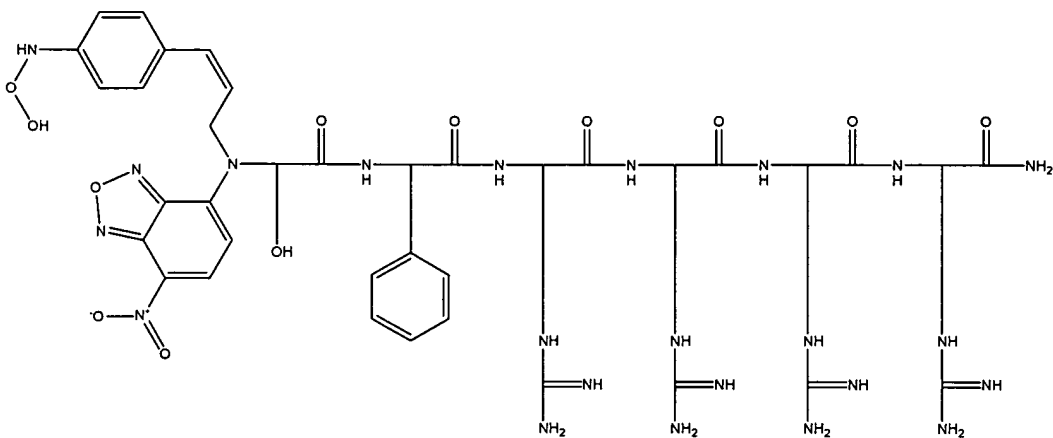


Compound 107

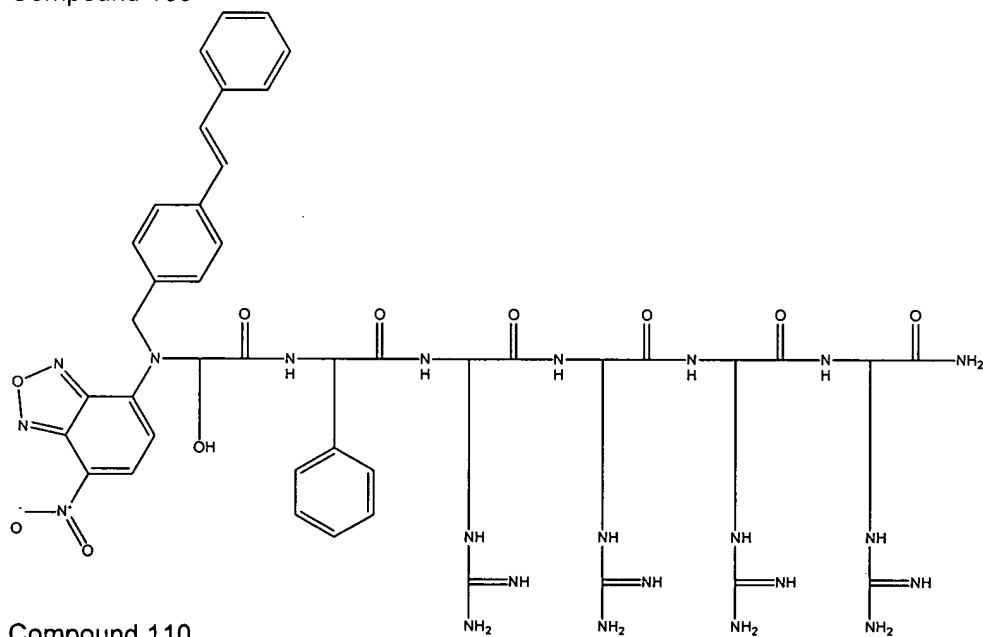


Compound 108

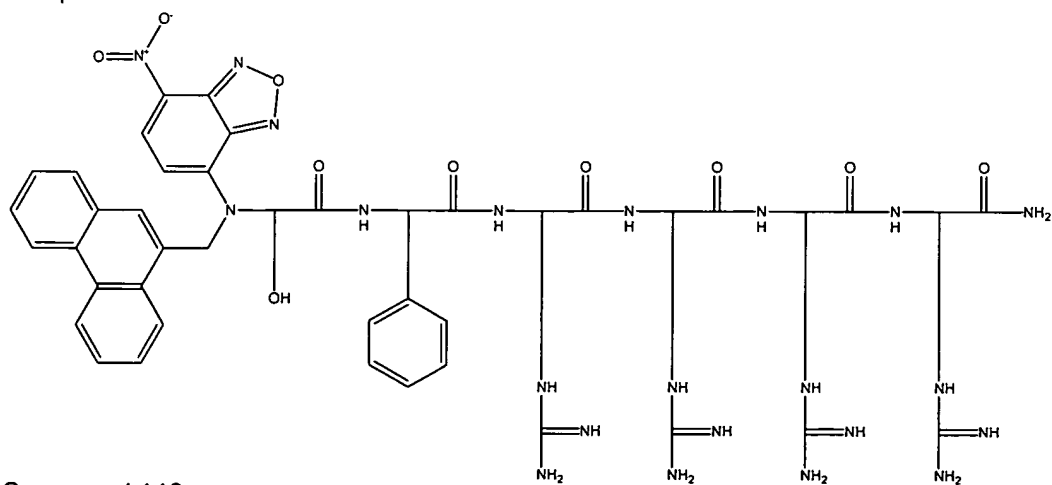
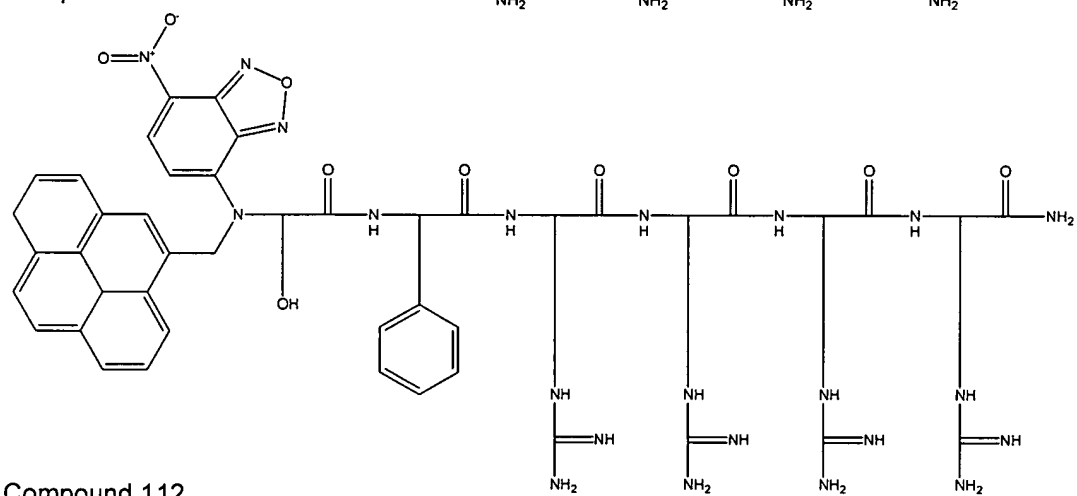
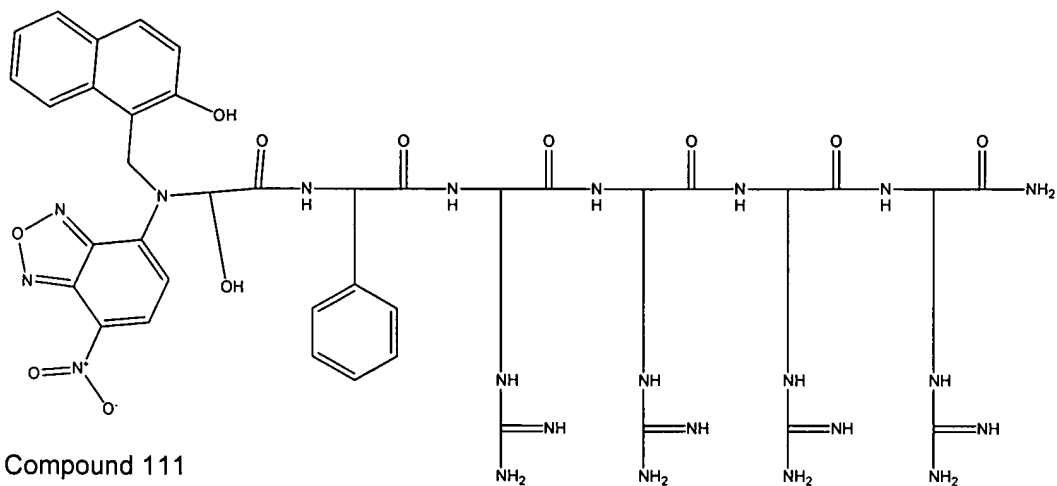
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 51 of 192

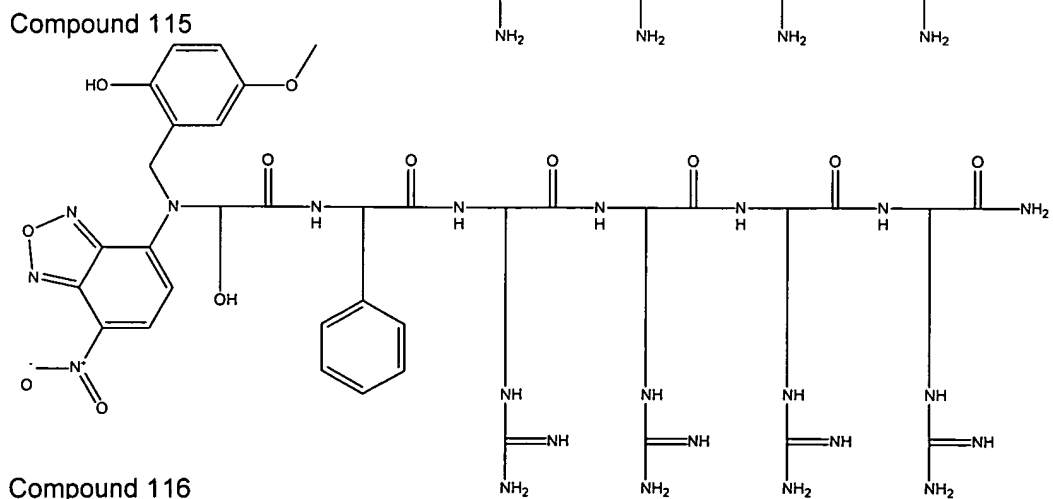
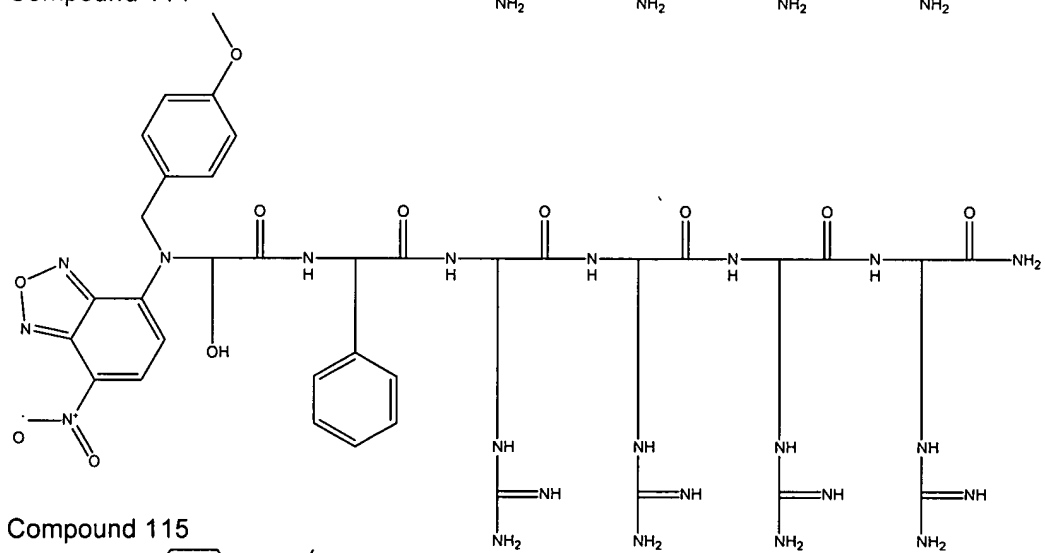
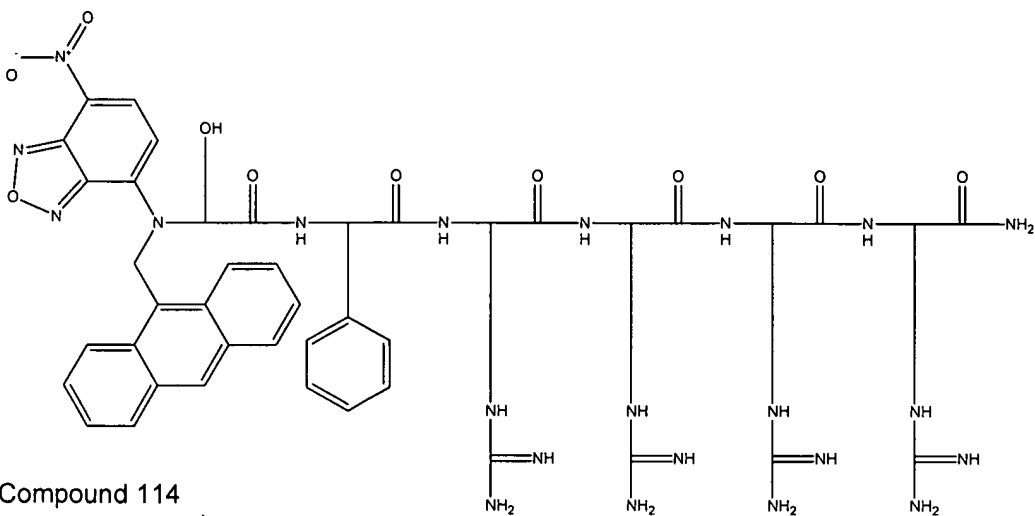


Compound 109

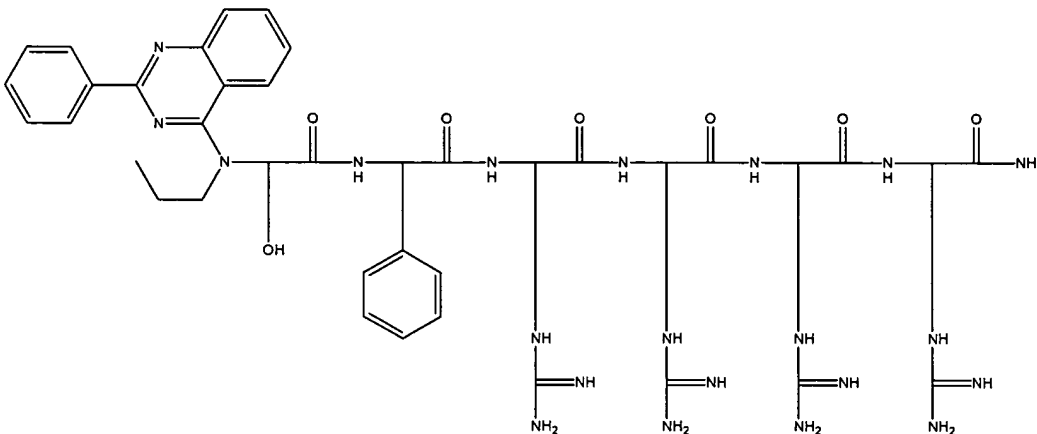


Compound 110

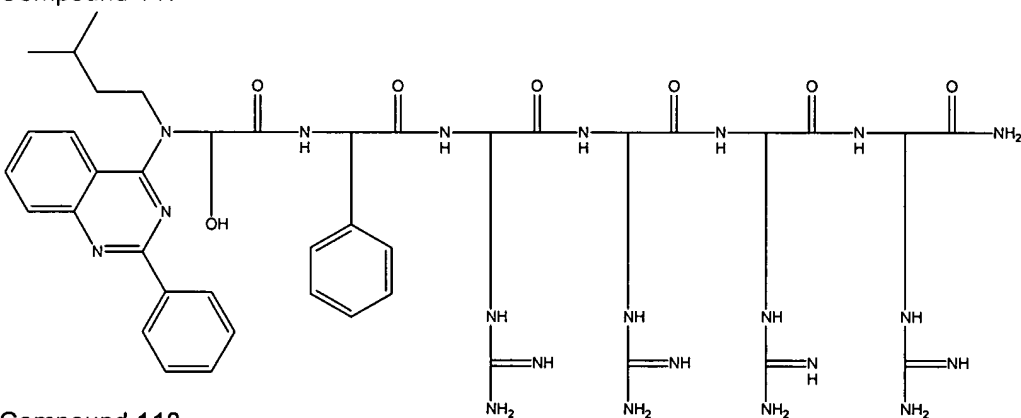




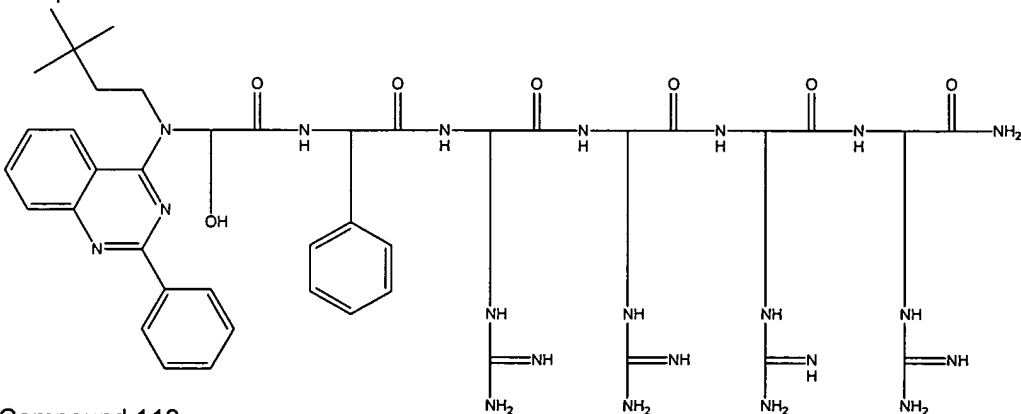
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 54 of 192



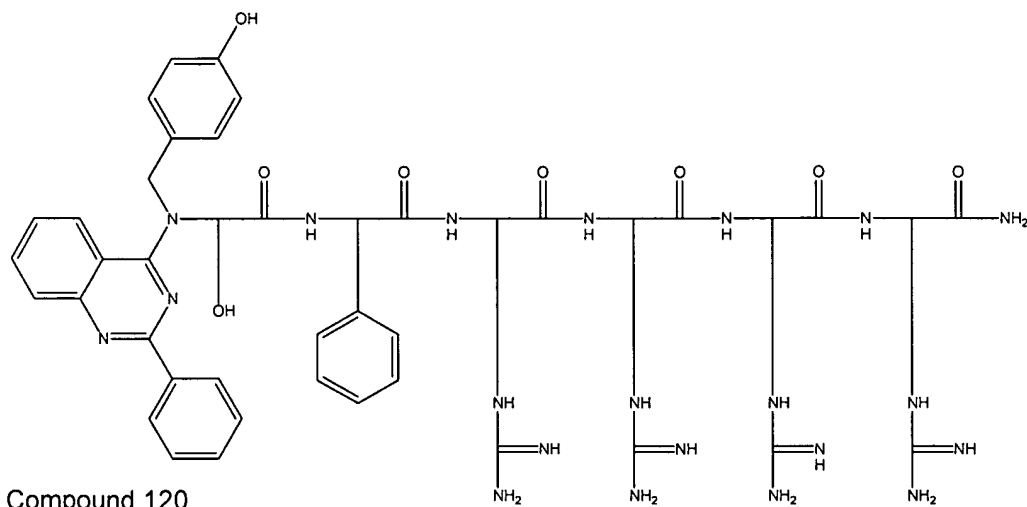
Compound 117



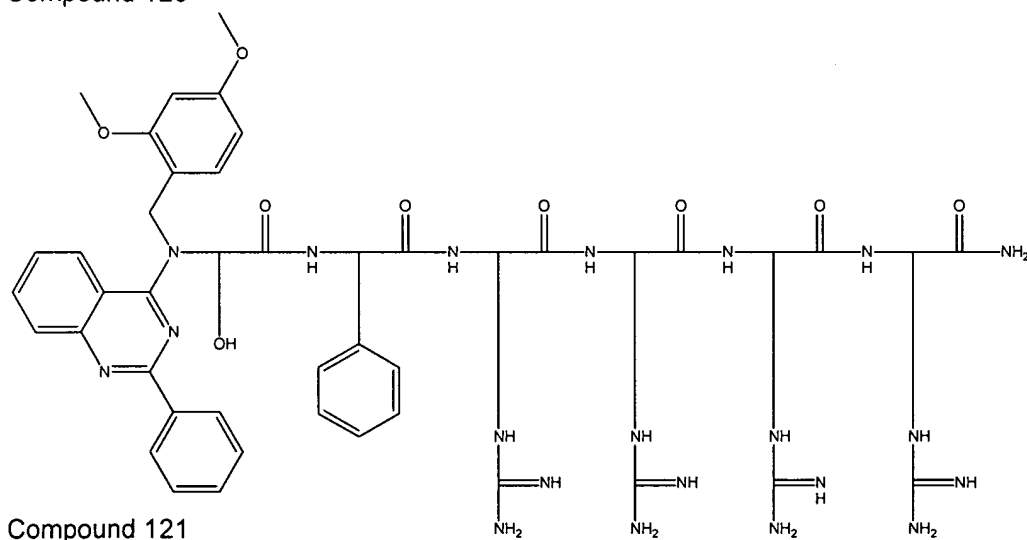
Compound 118



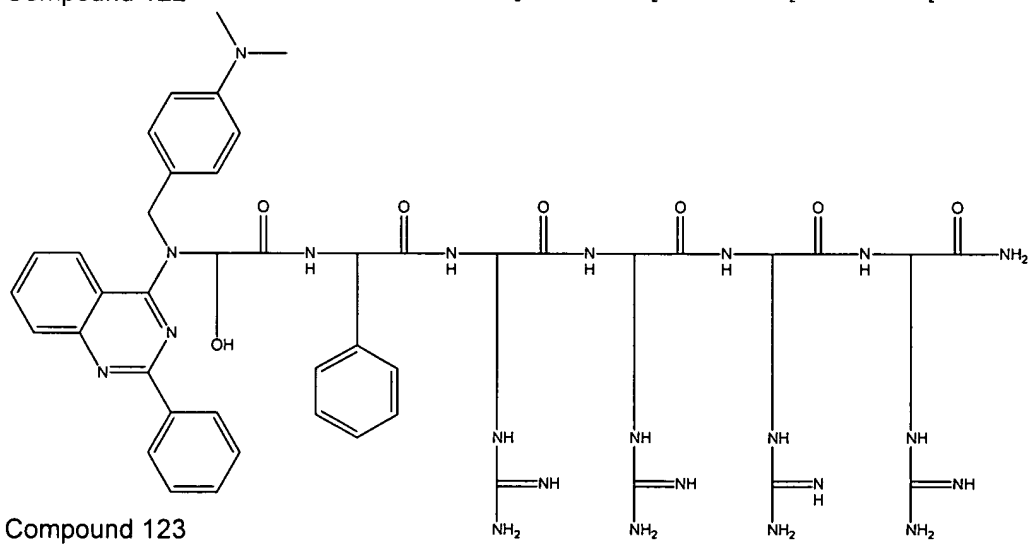
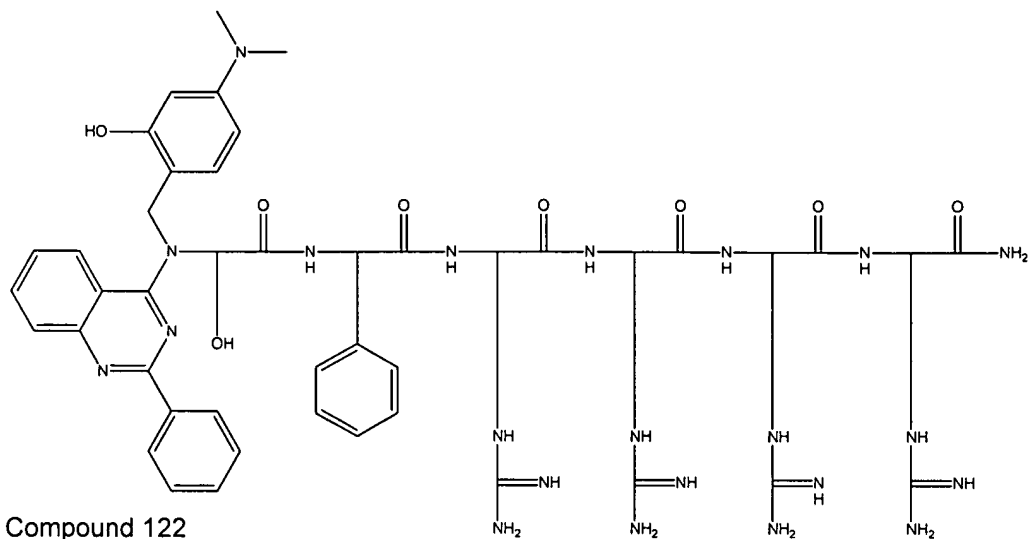
Compound 119

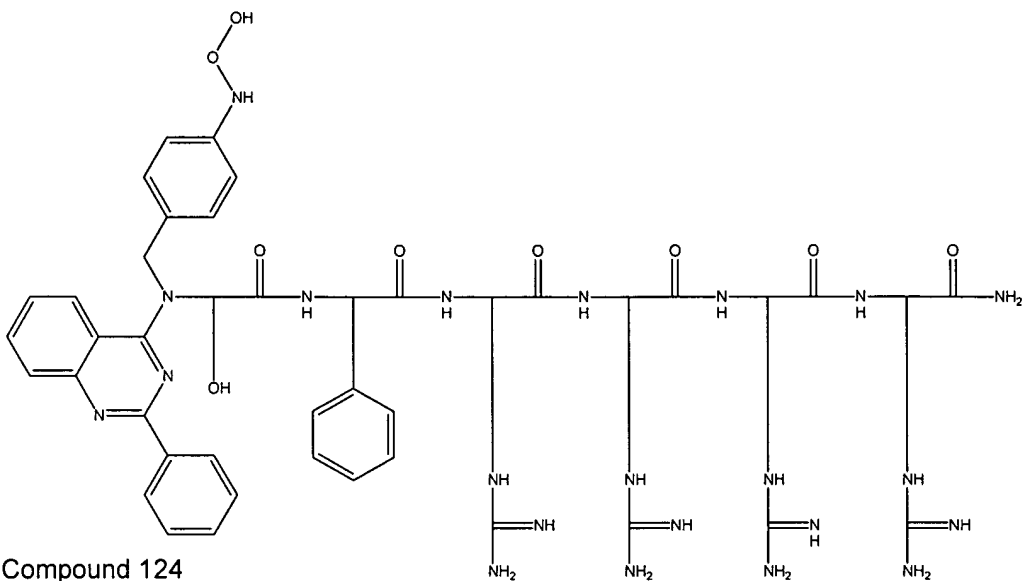


Compound 120

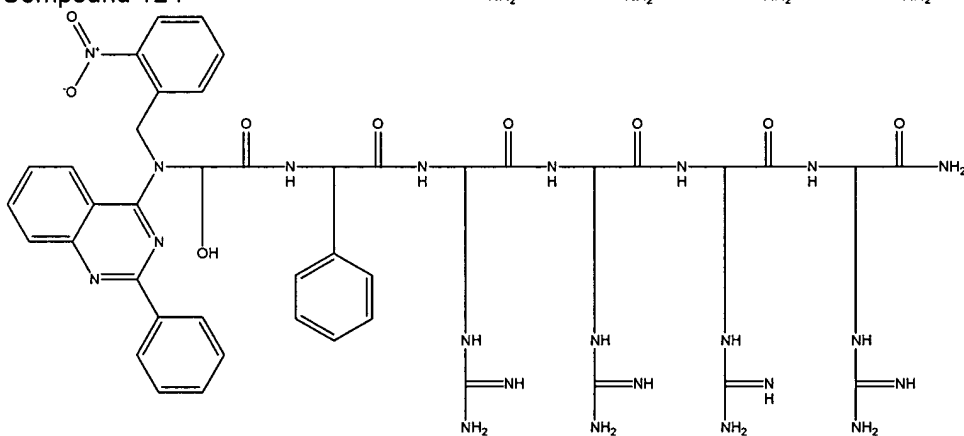


Compound 121



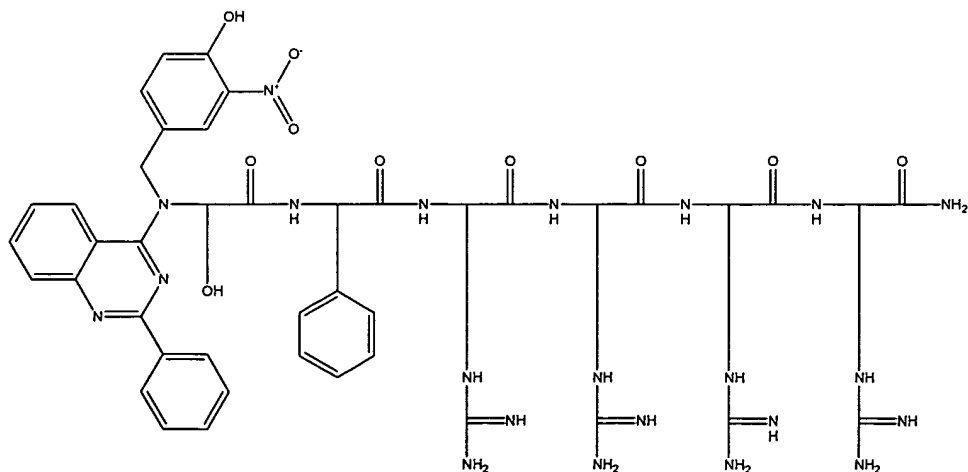


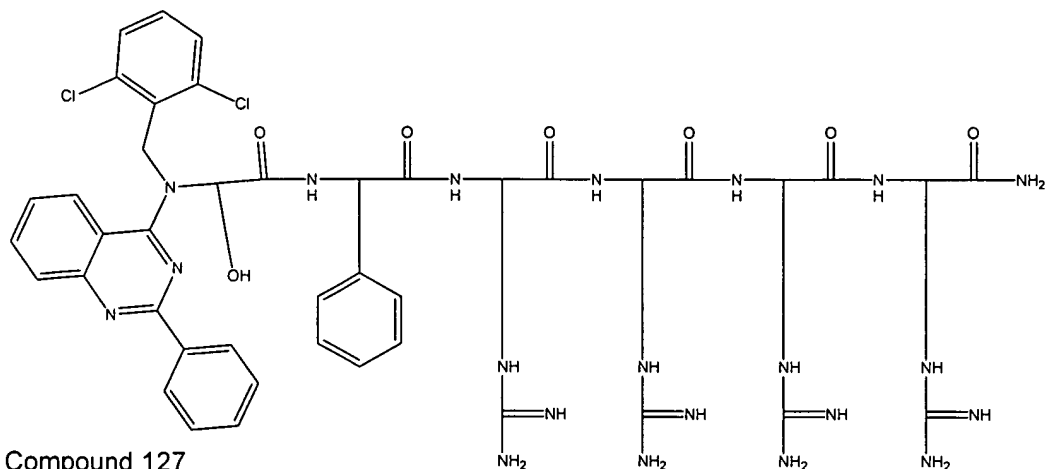
Compound 124



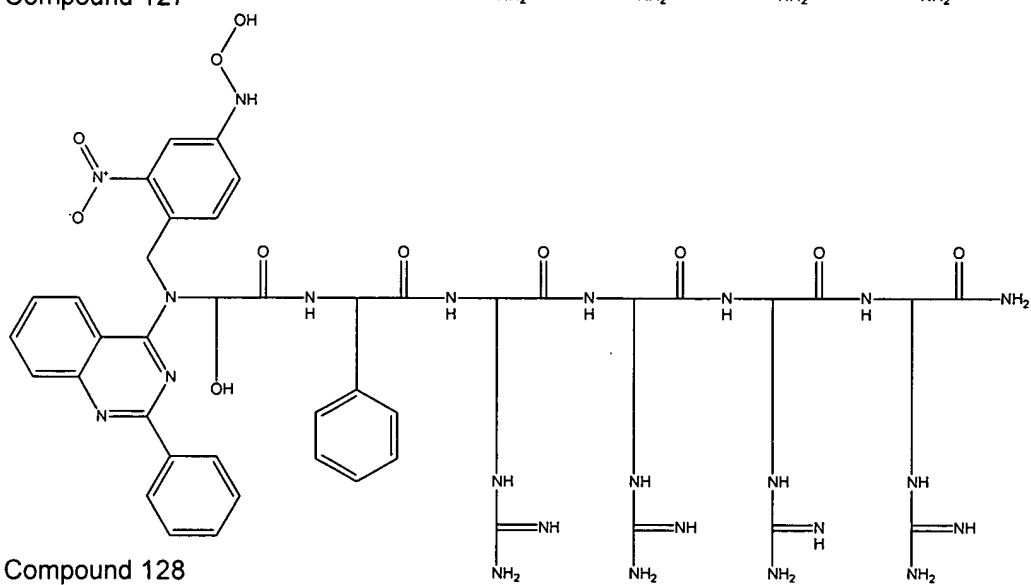
Compound 125

Compound 126

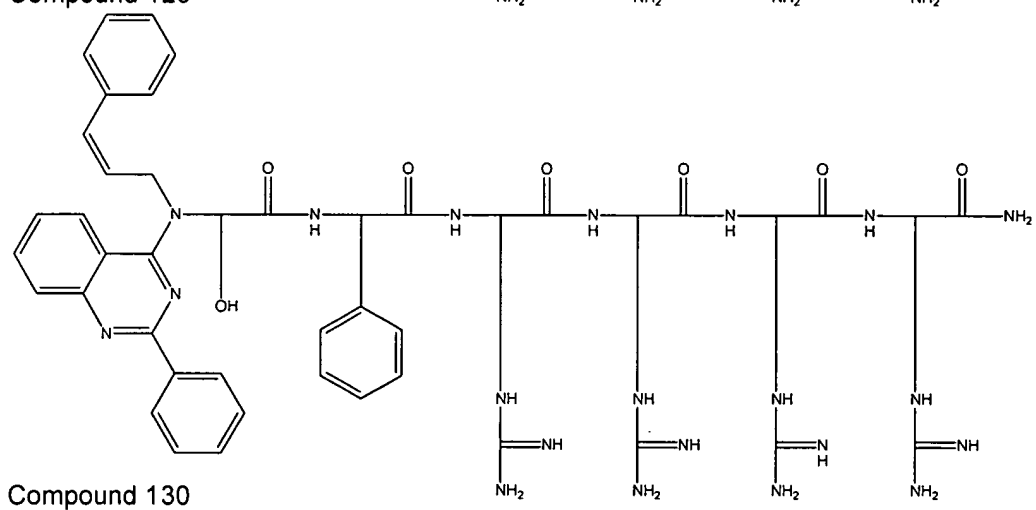
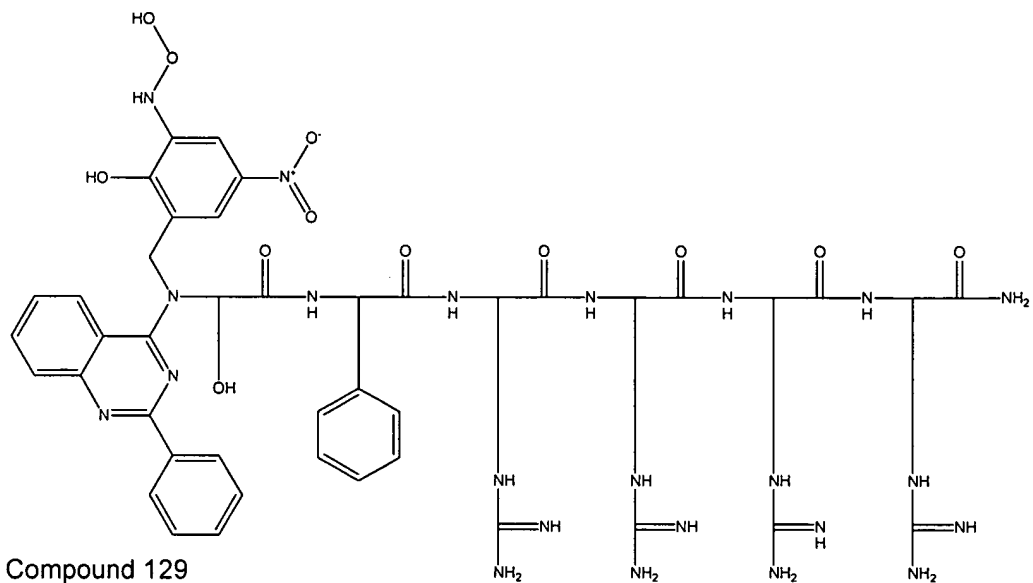




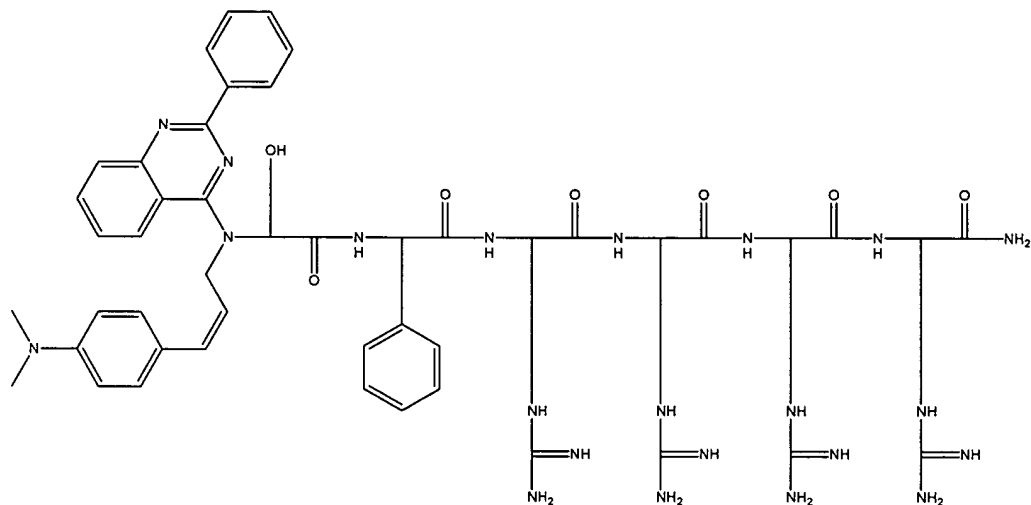
Compound 127



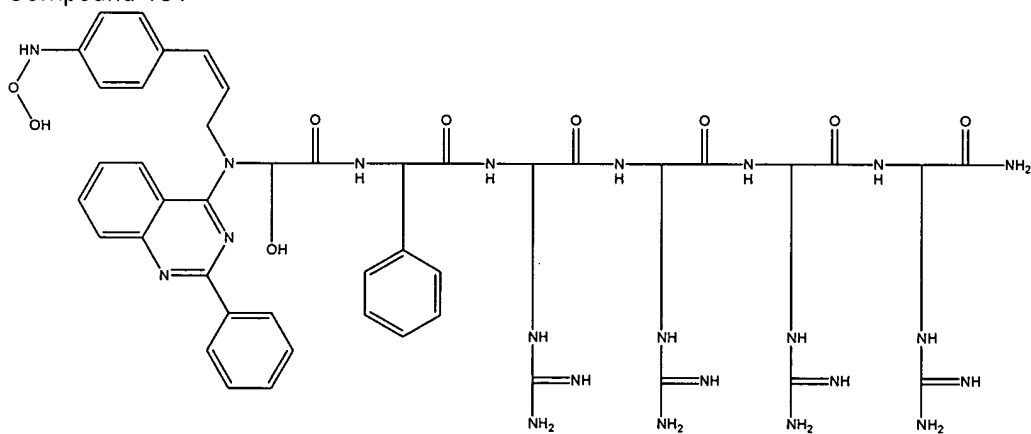
Compound 128



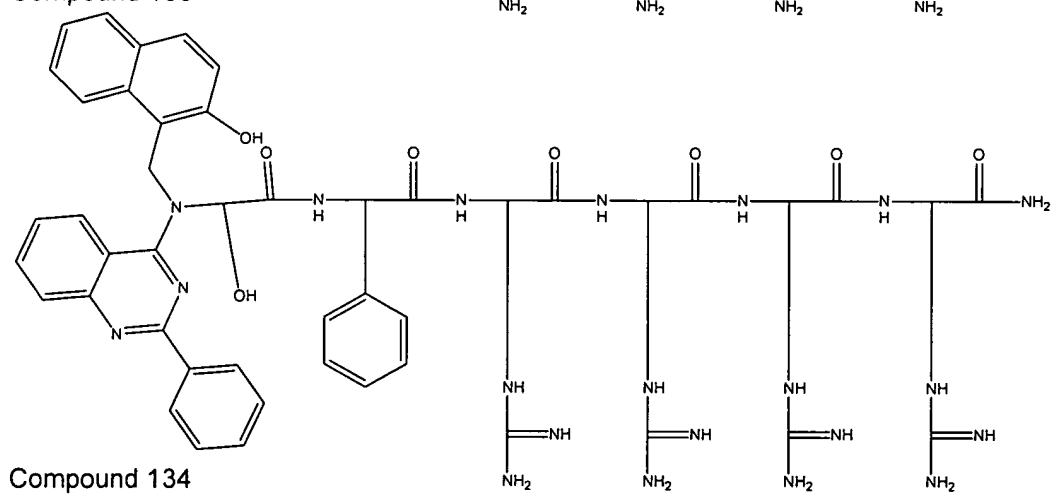
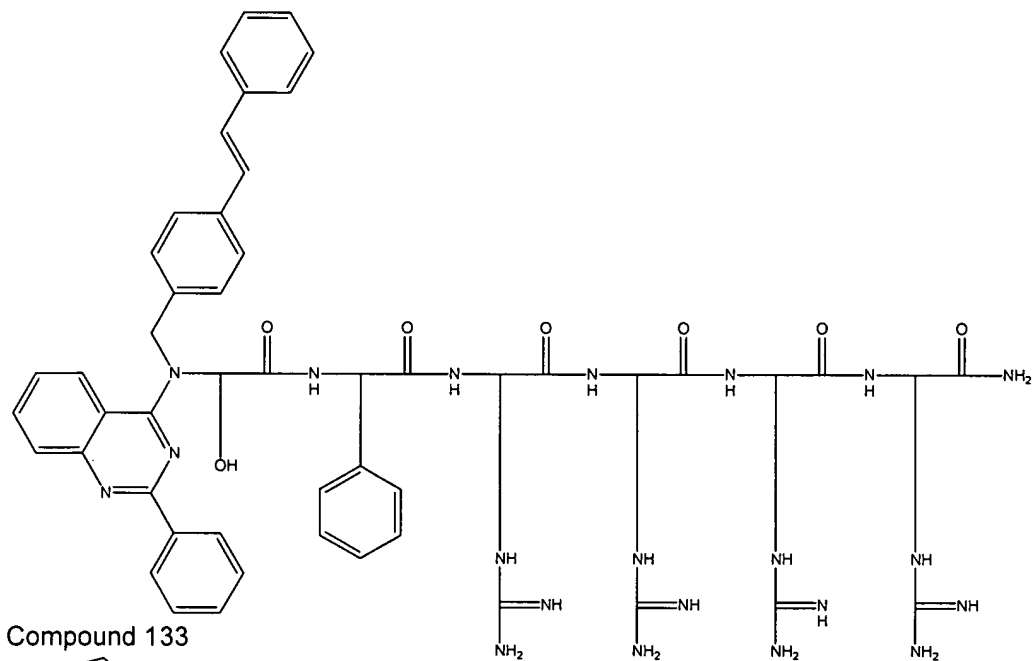
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 60 of 192



Compound 131

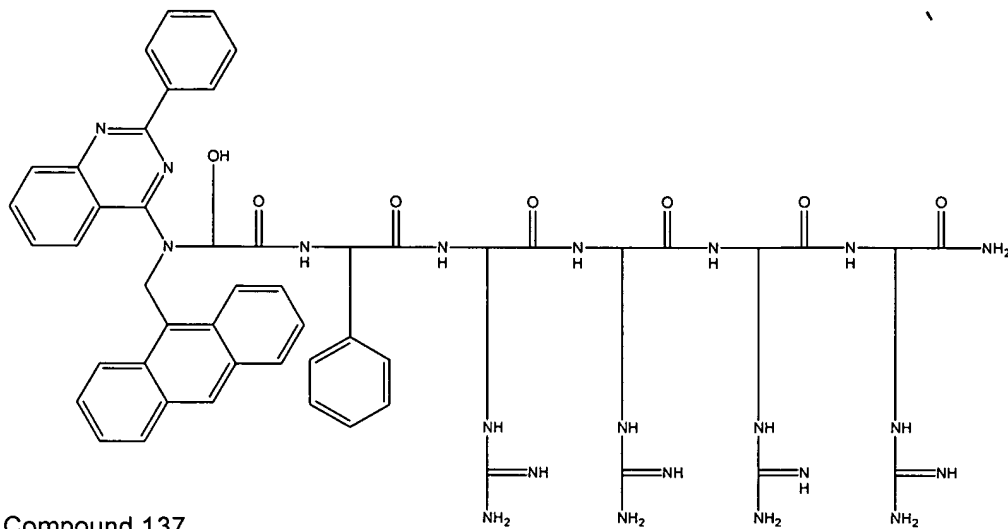


Compound 132

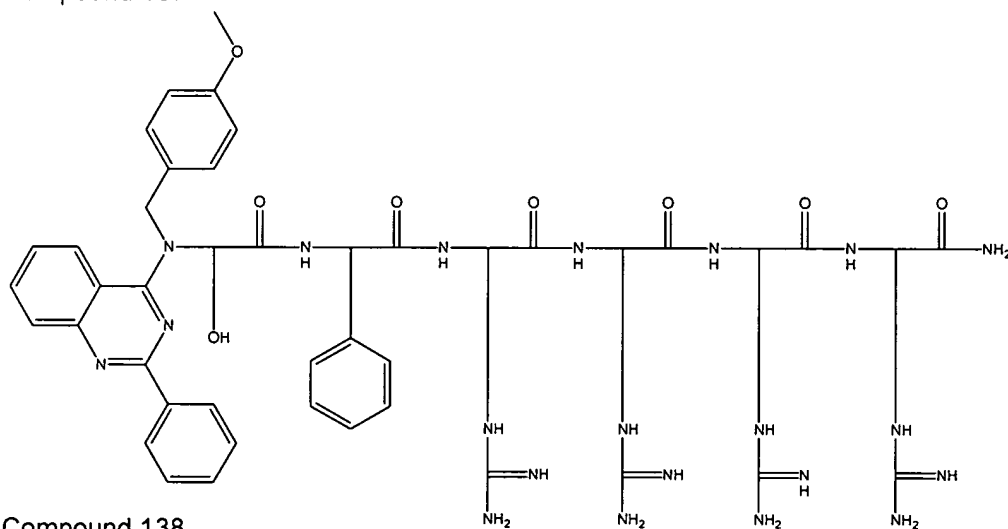


page 62 of 192

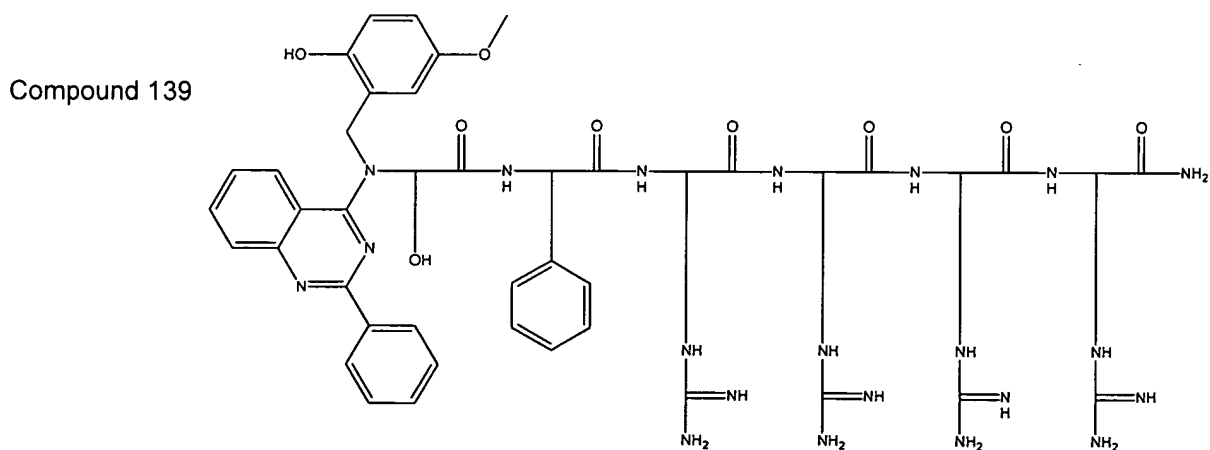




Compound 137



Compound 138

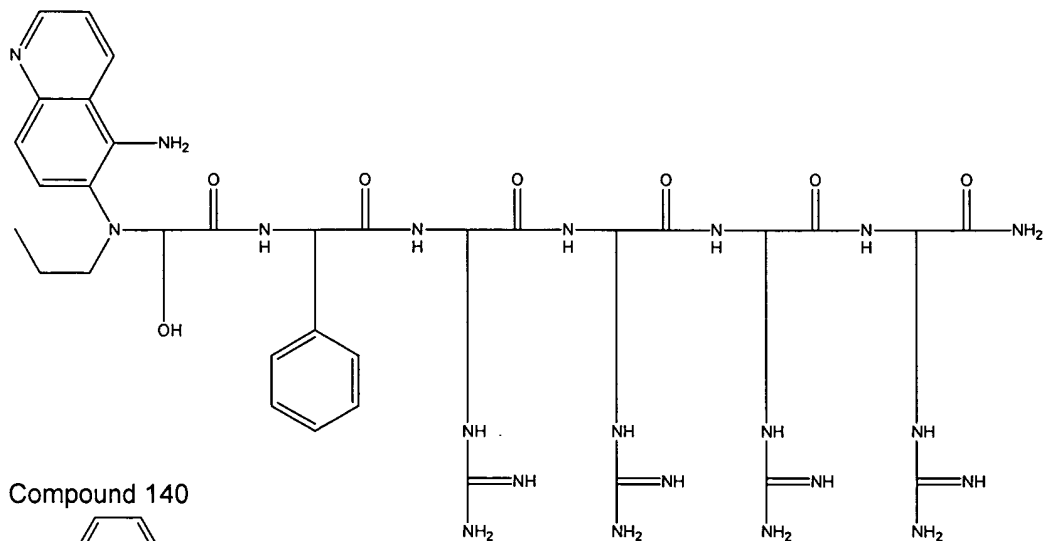


Applicant: David S. Lawrence

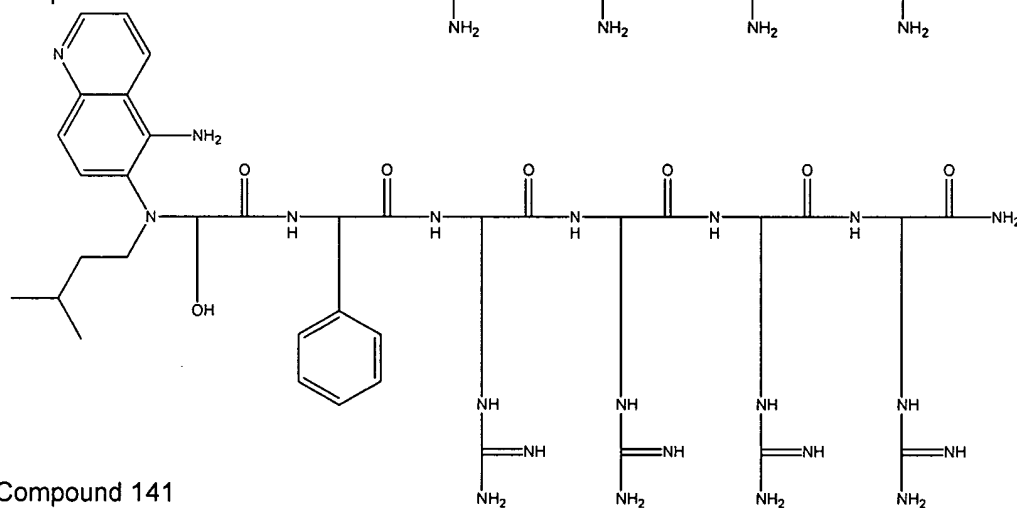
Serial No.: 10/755,086

Filed: January 9, 2004

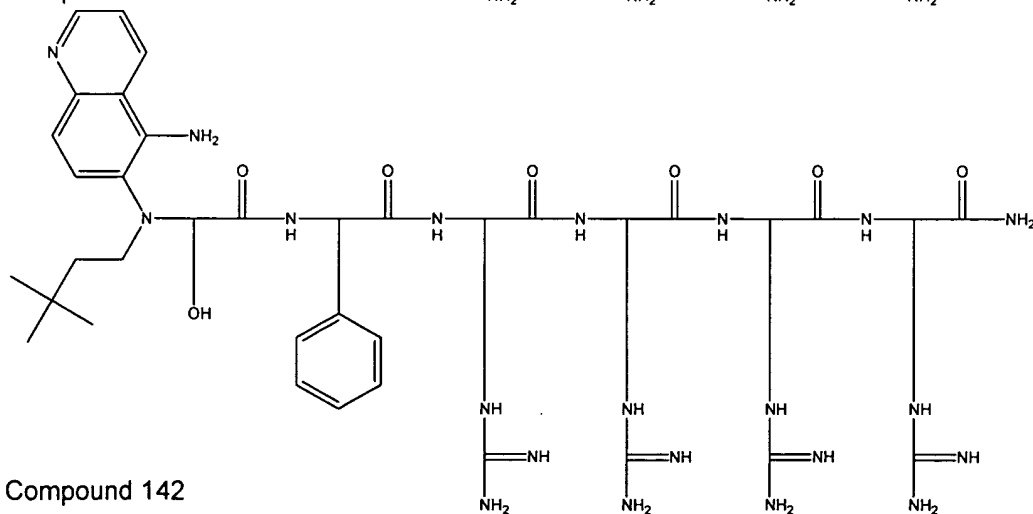
page 64 of 192



Compound 140



Compound 141



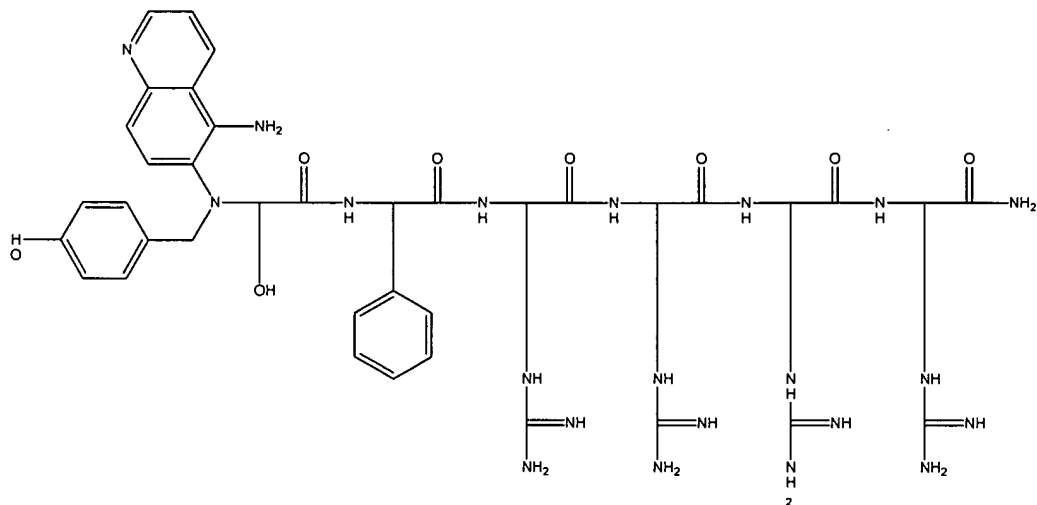
Compound 142

Applicant: David S. Lawrence

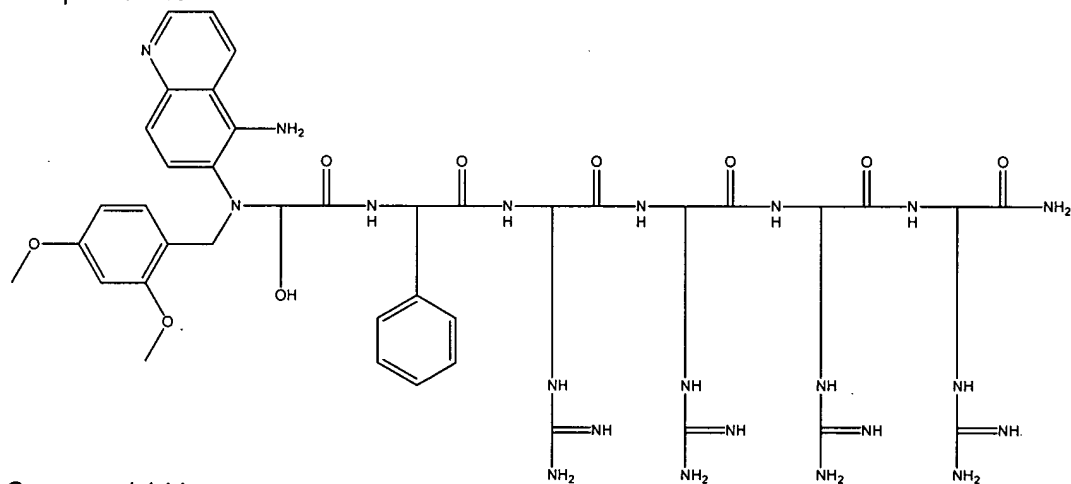
Serial No.: 10/755,086

Filed: January 9, 2004

page 65 of 192

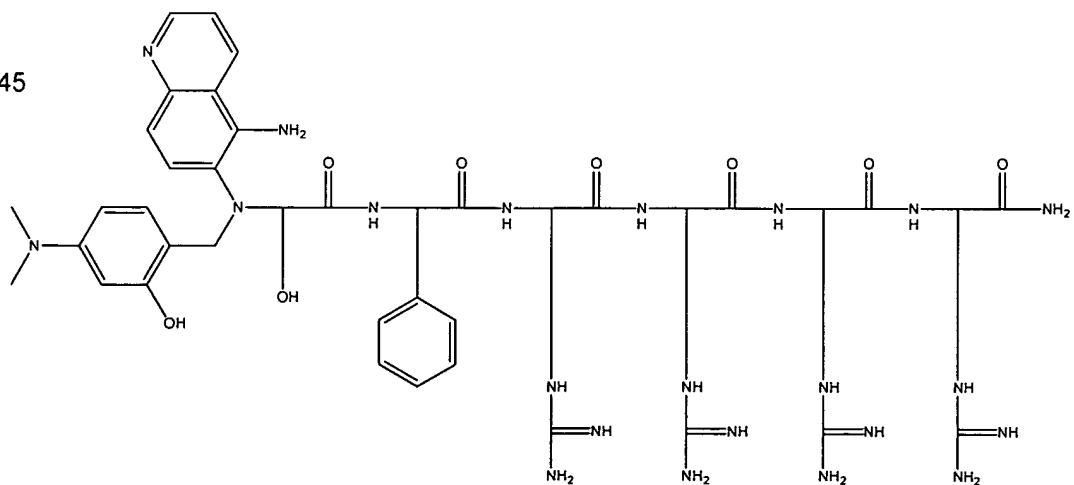


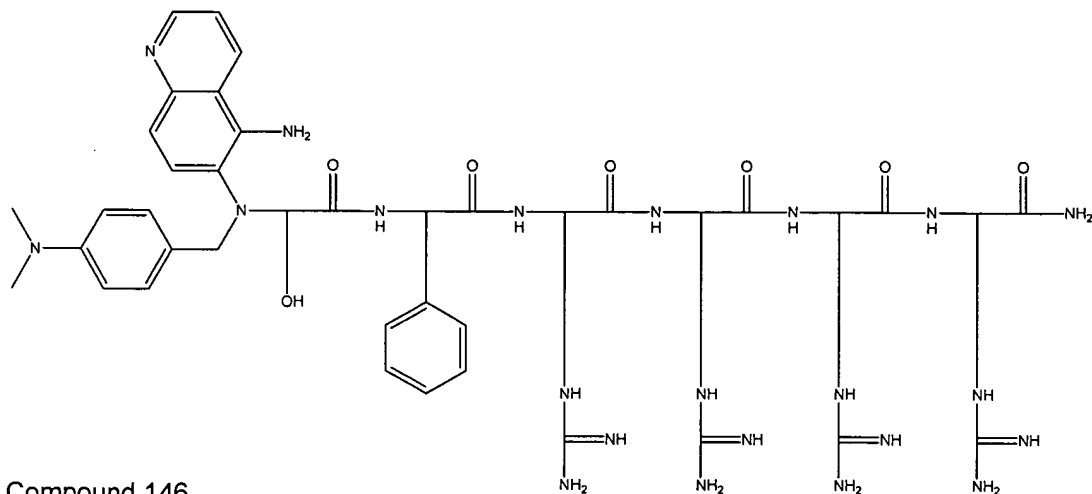
Compound 143



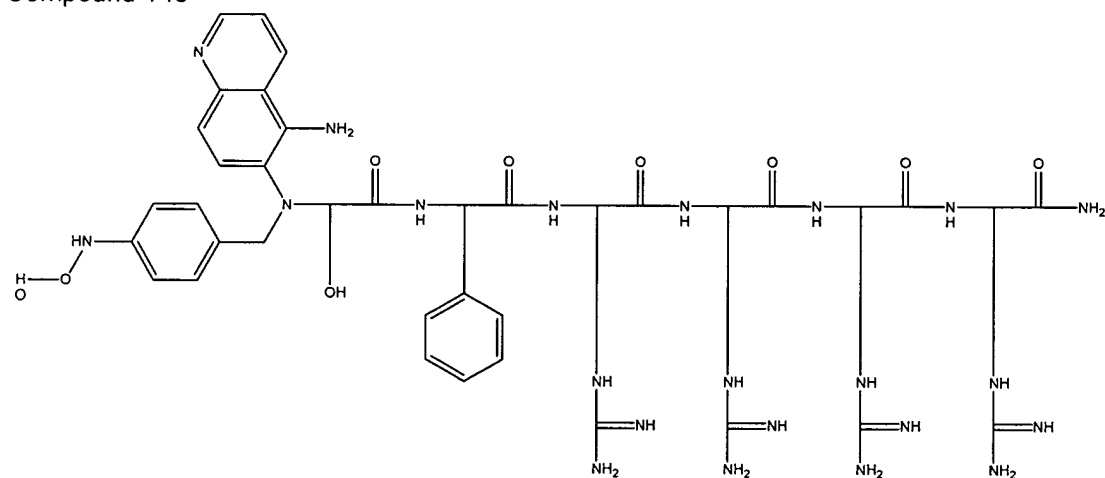
Compound 144

Compound 145

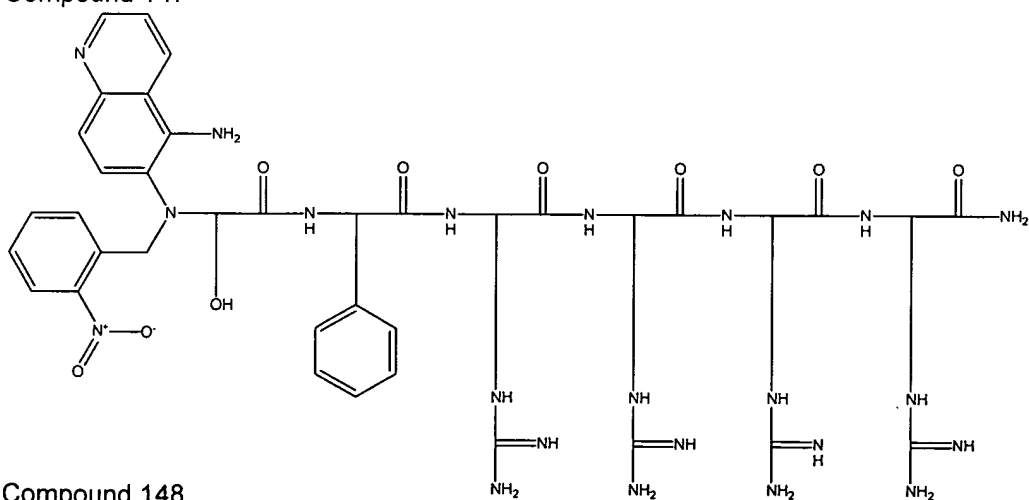




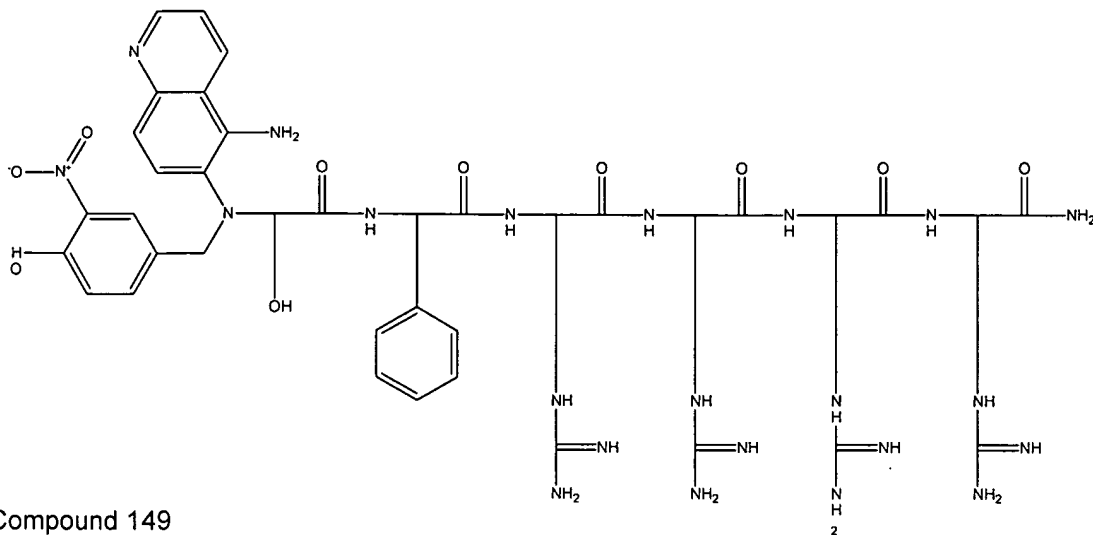
Compound 146



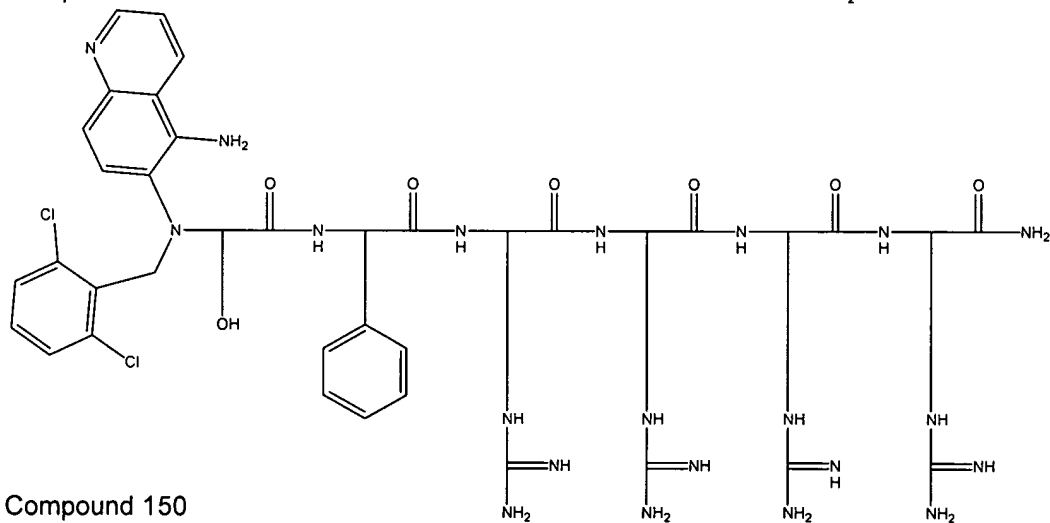
Compound 147



Compound 148

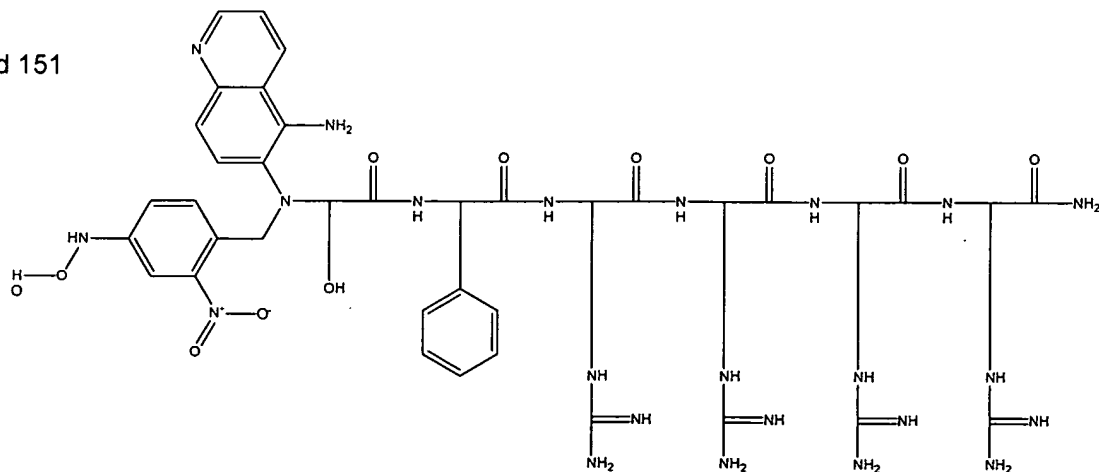


Compound 149



Compound 150

Compound 151

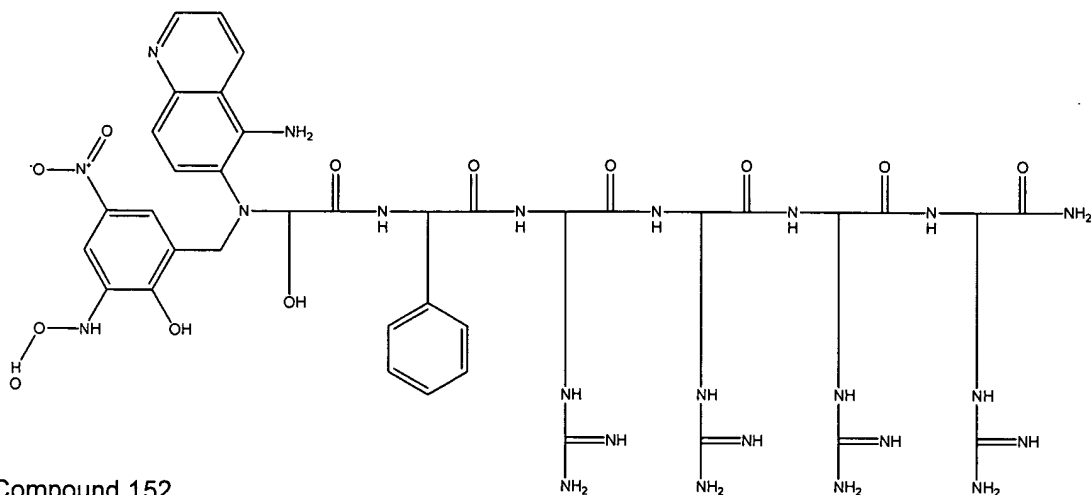


Applicant: David S. Lawrence

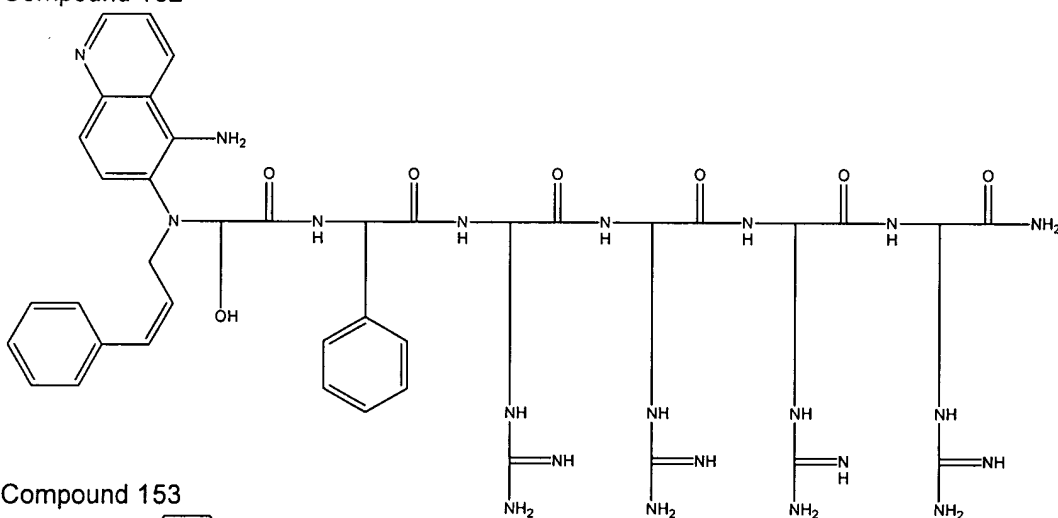
Serial No.: 10/755,086

Filed: January 9, 2004

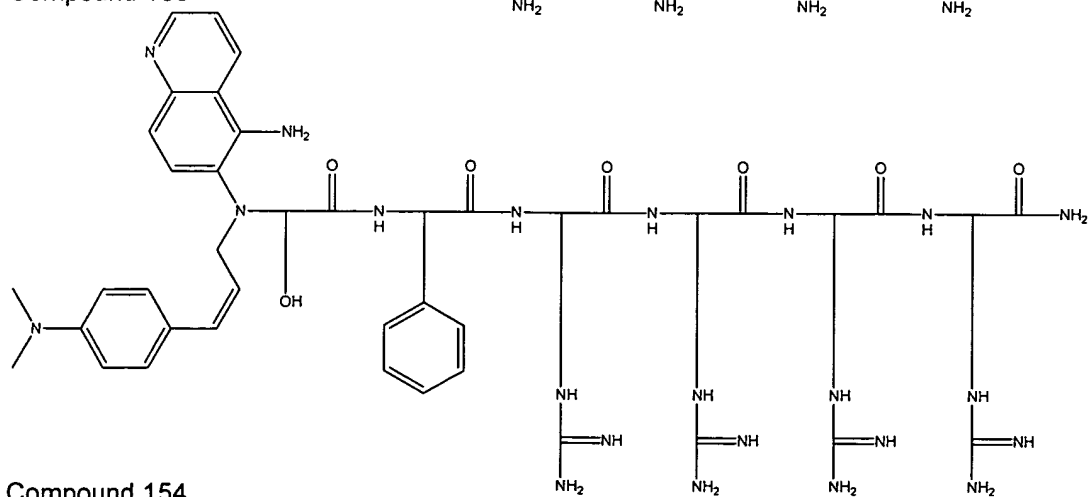
page 68 of 192



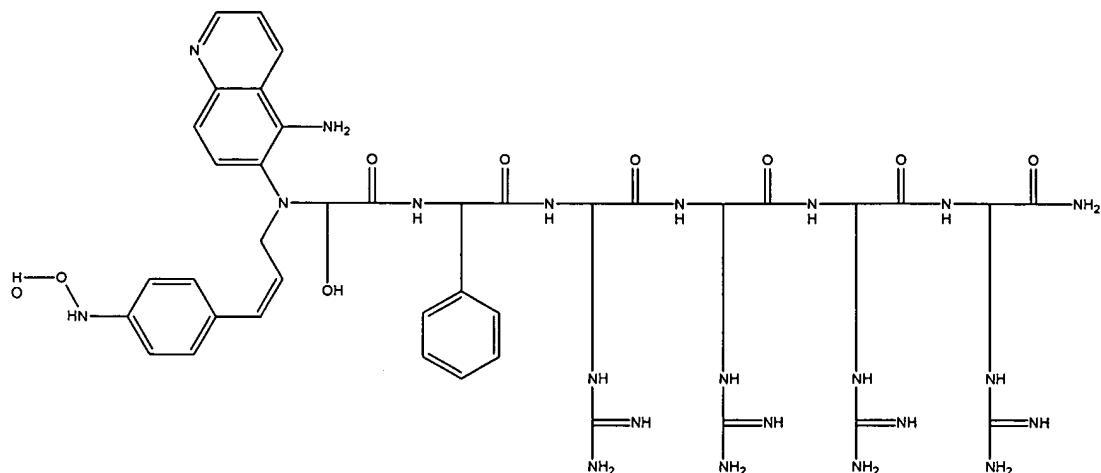
Compound 152



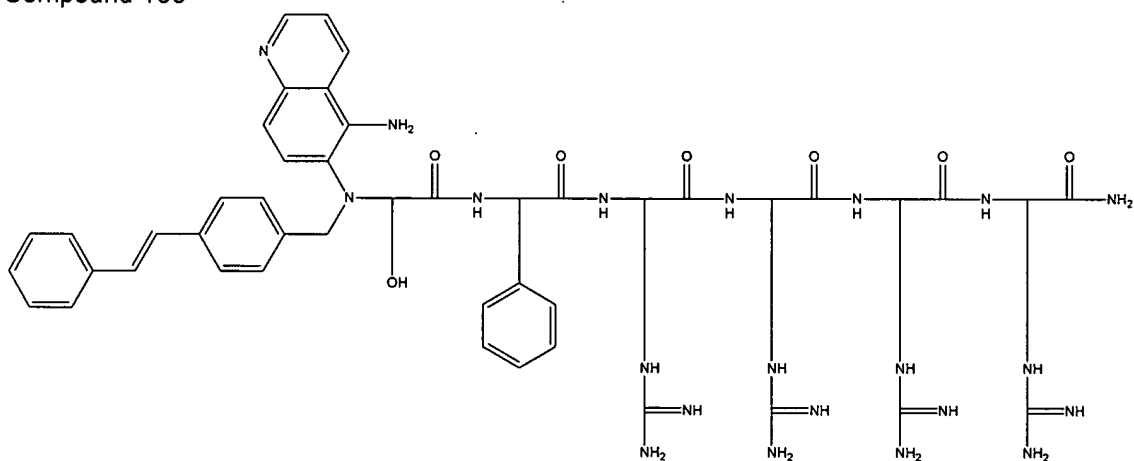
Compound 153



Compound 154

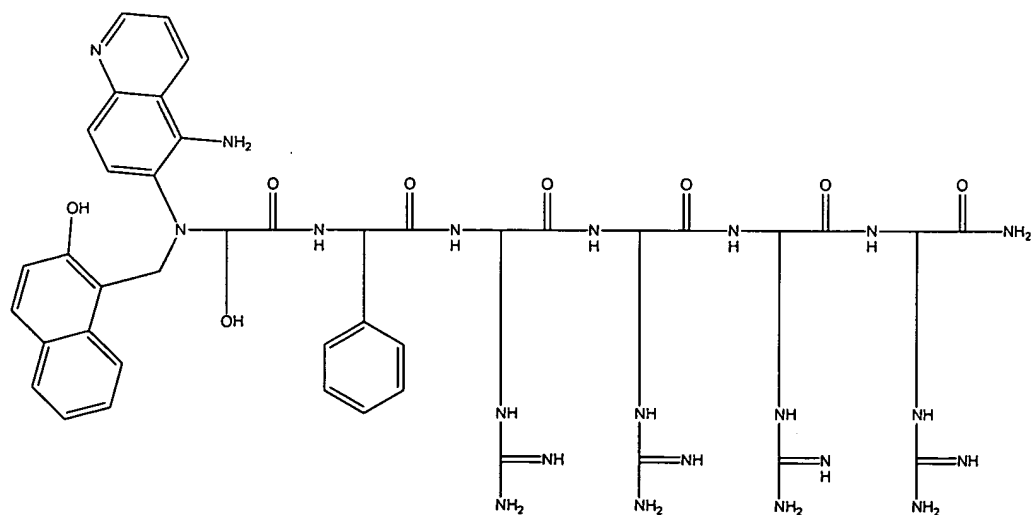


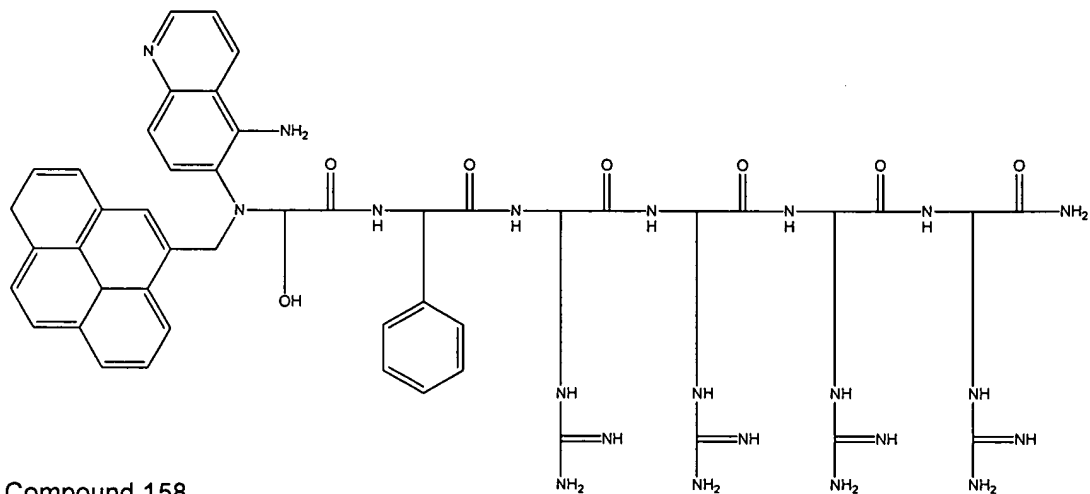
Compound 155



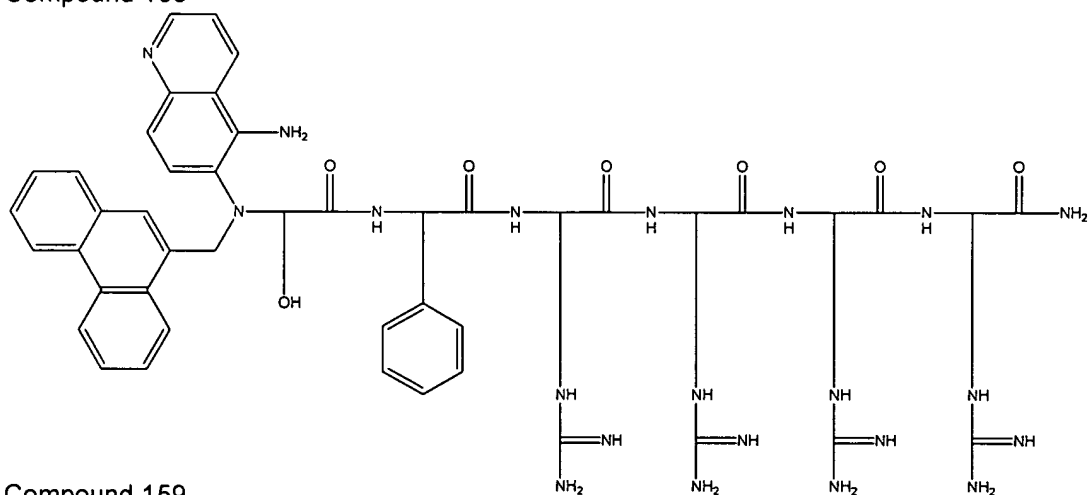
Compound 156

Compound 157

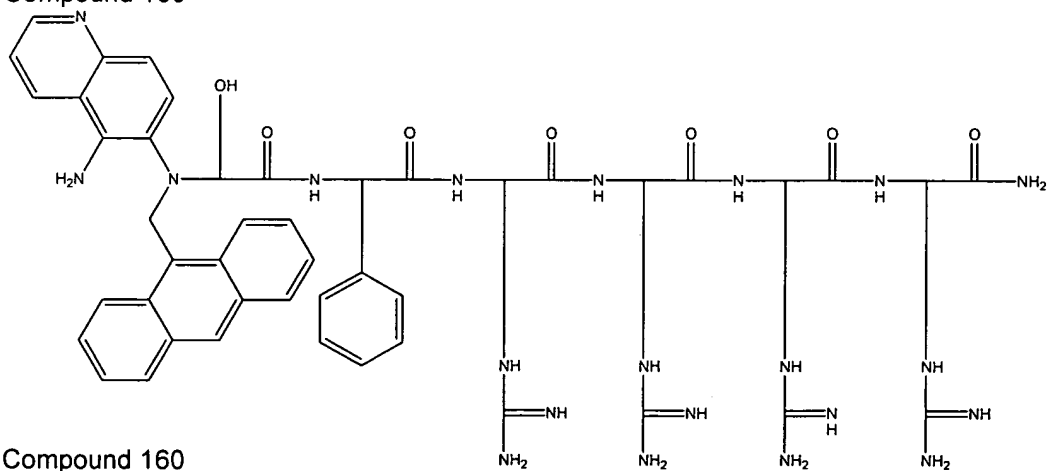




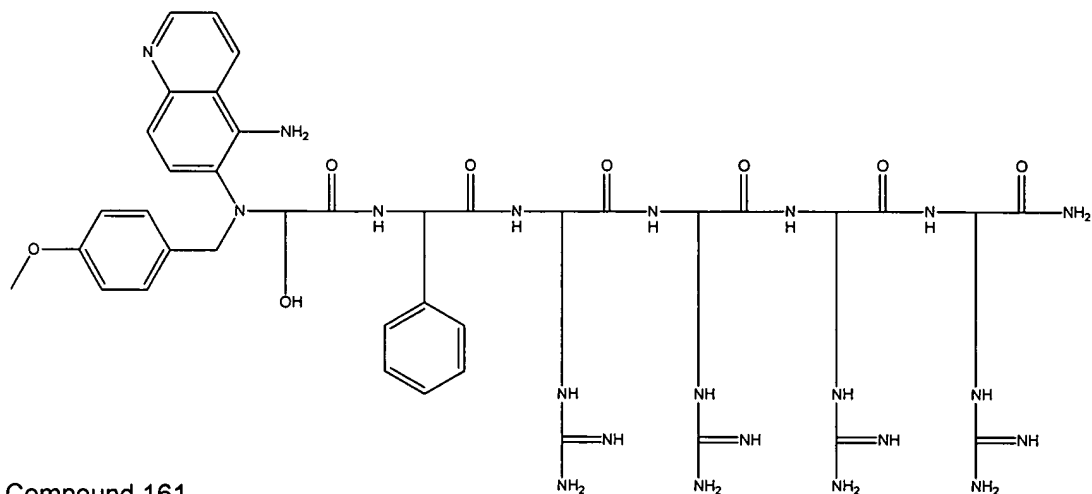
Compound 158



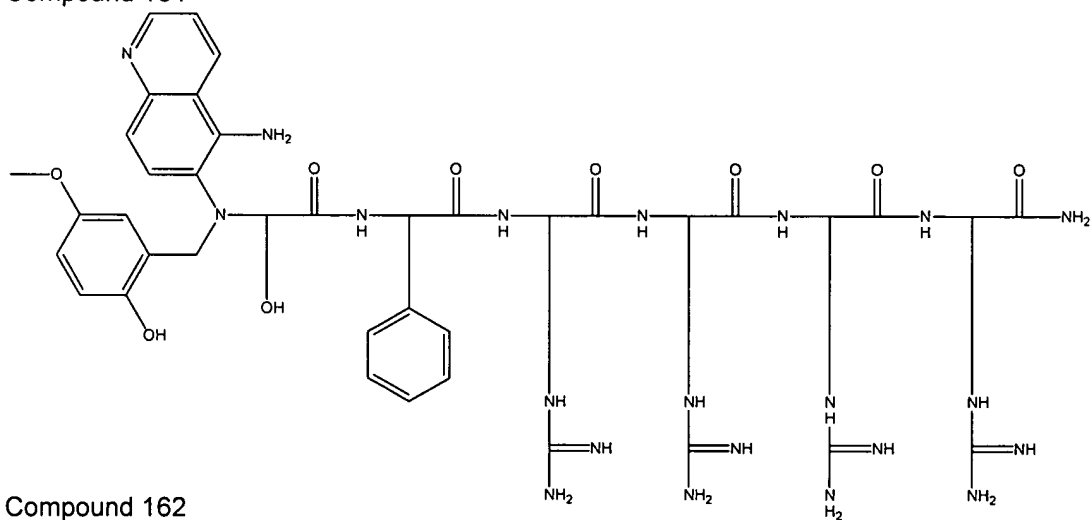
Compound 159



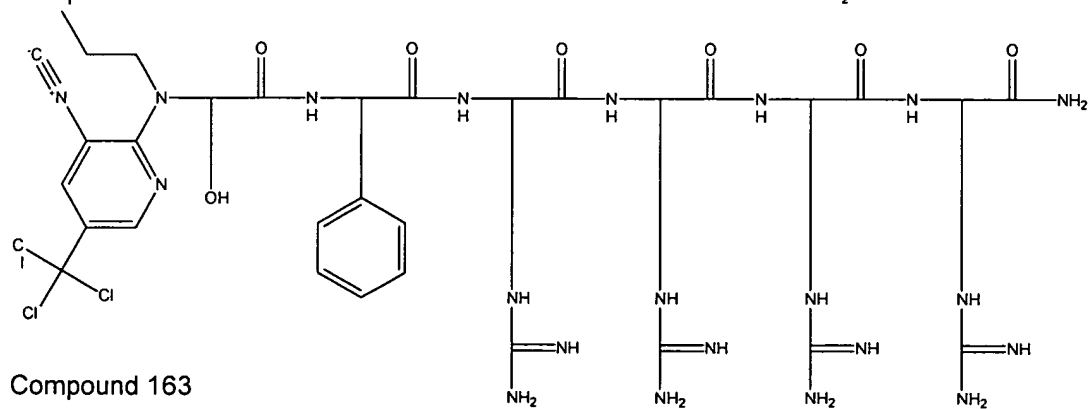
Compound 160



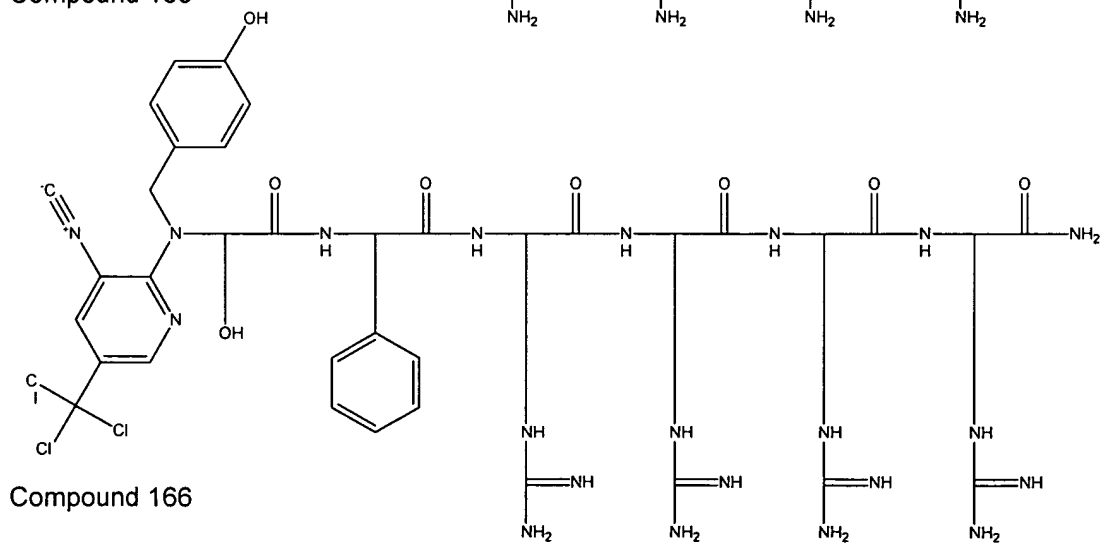
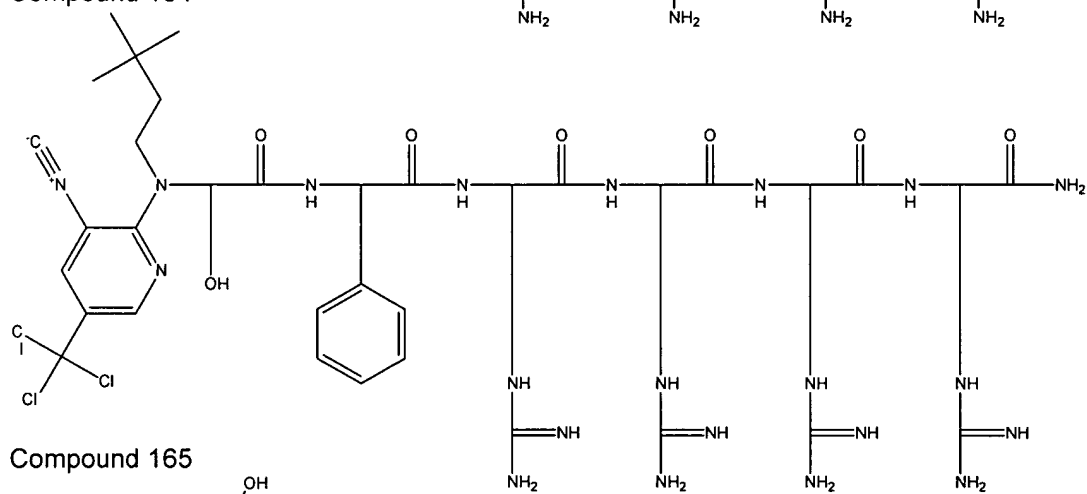
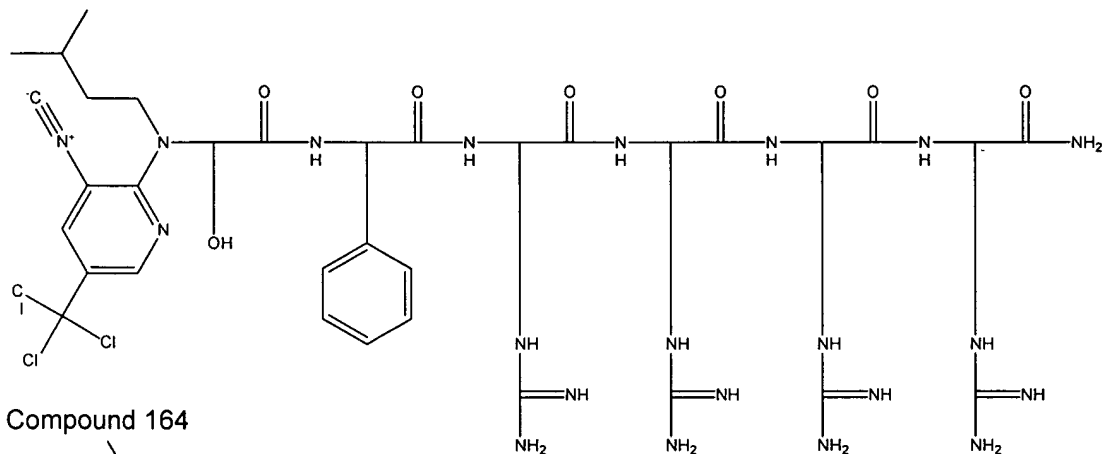
Compound 161

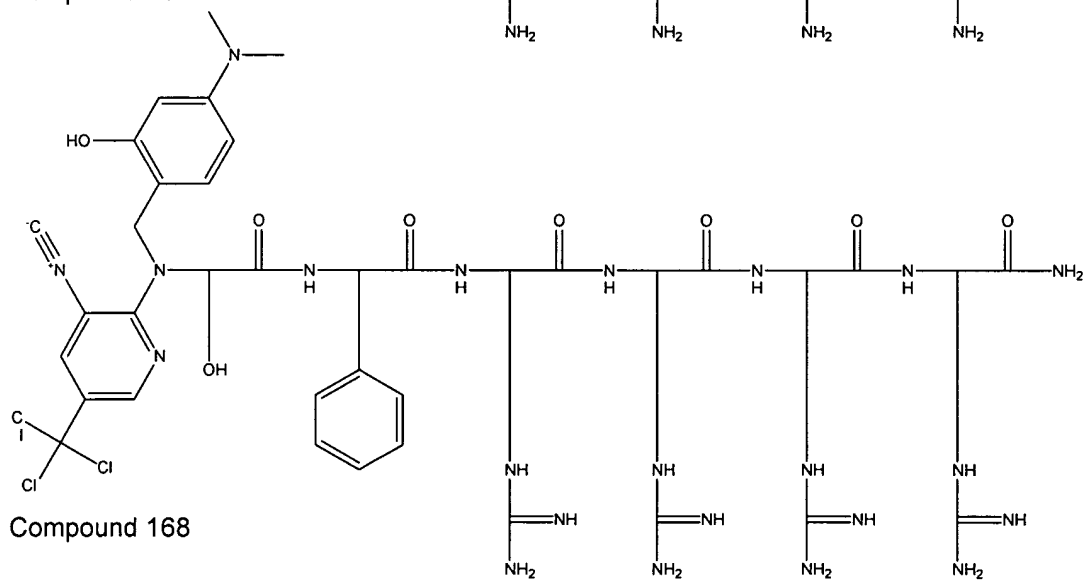
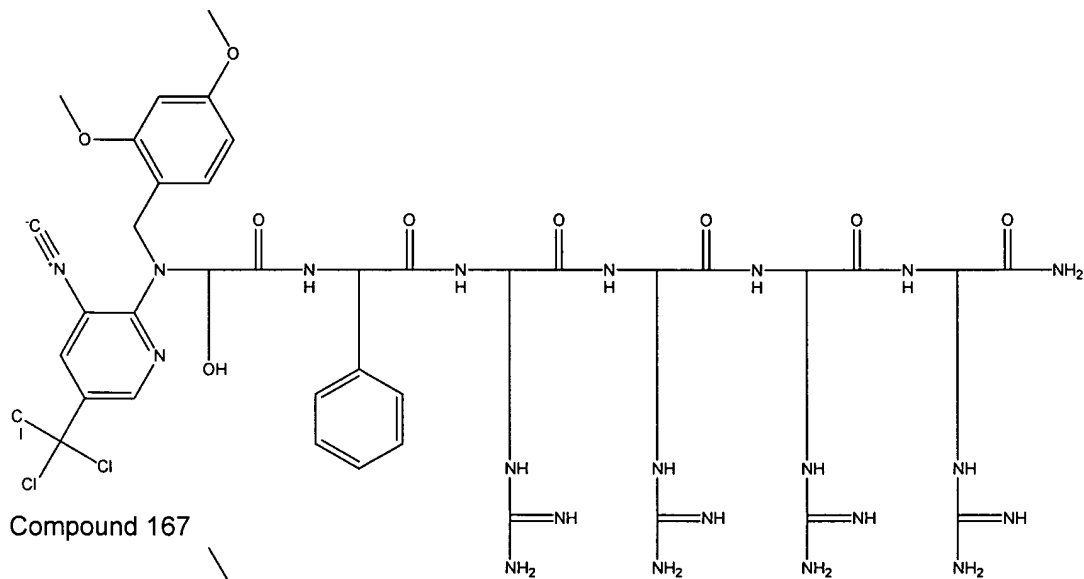


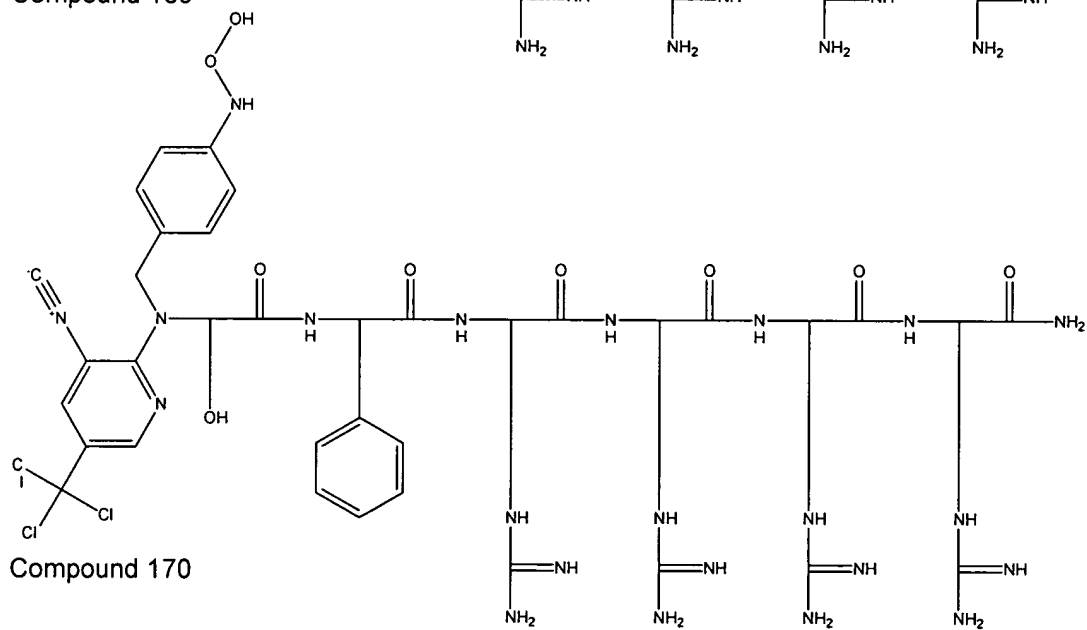
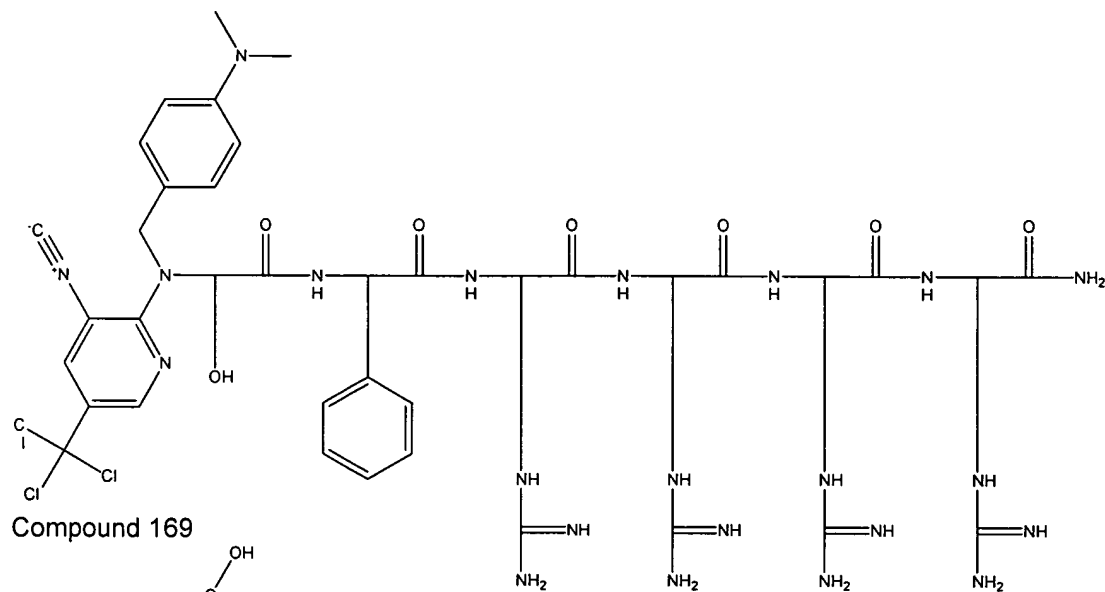
Compound 162

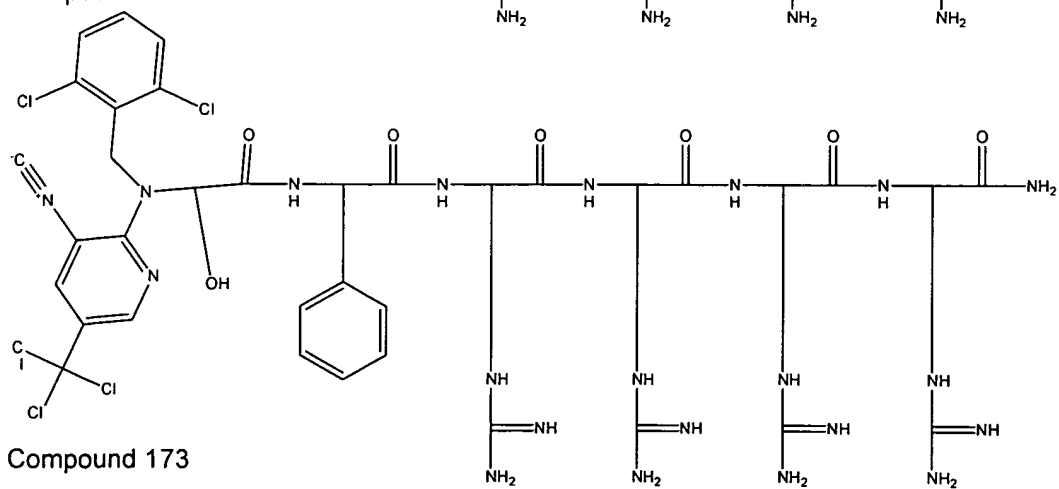
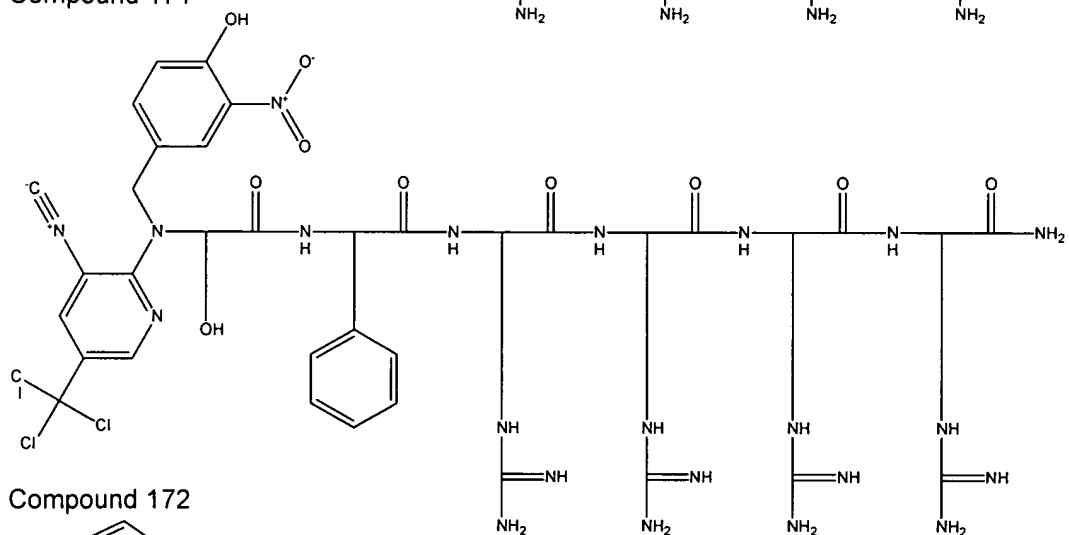
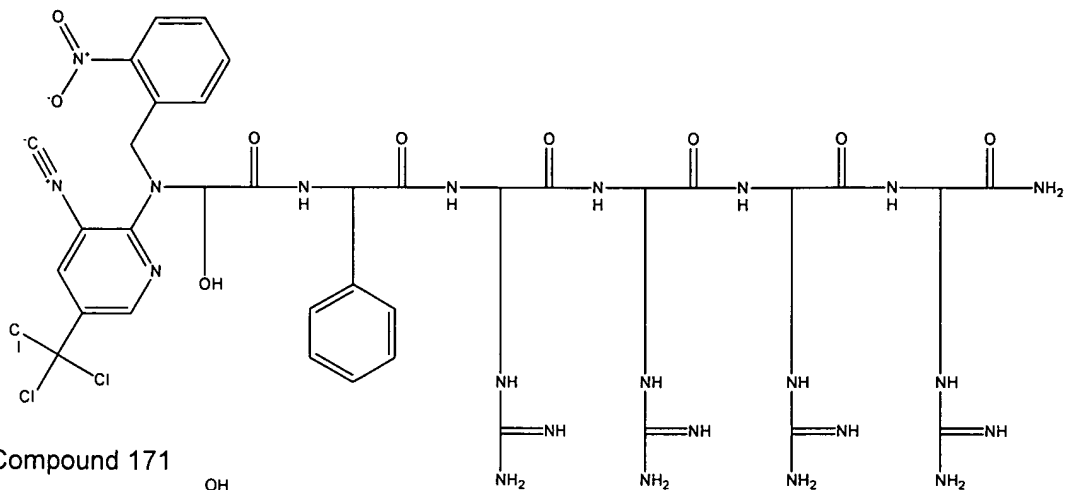


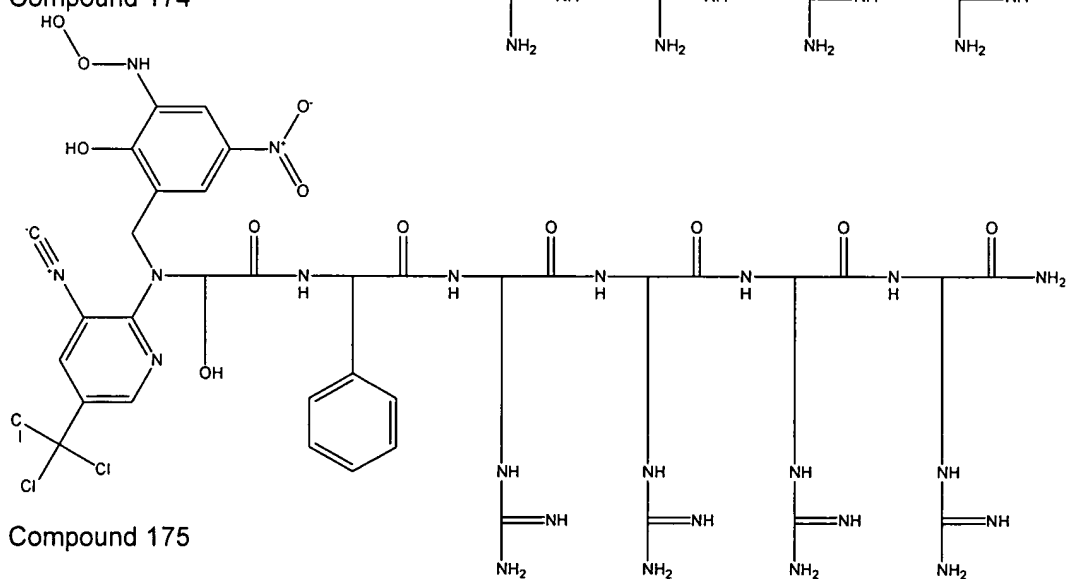
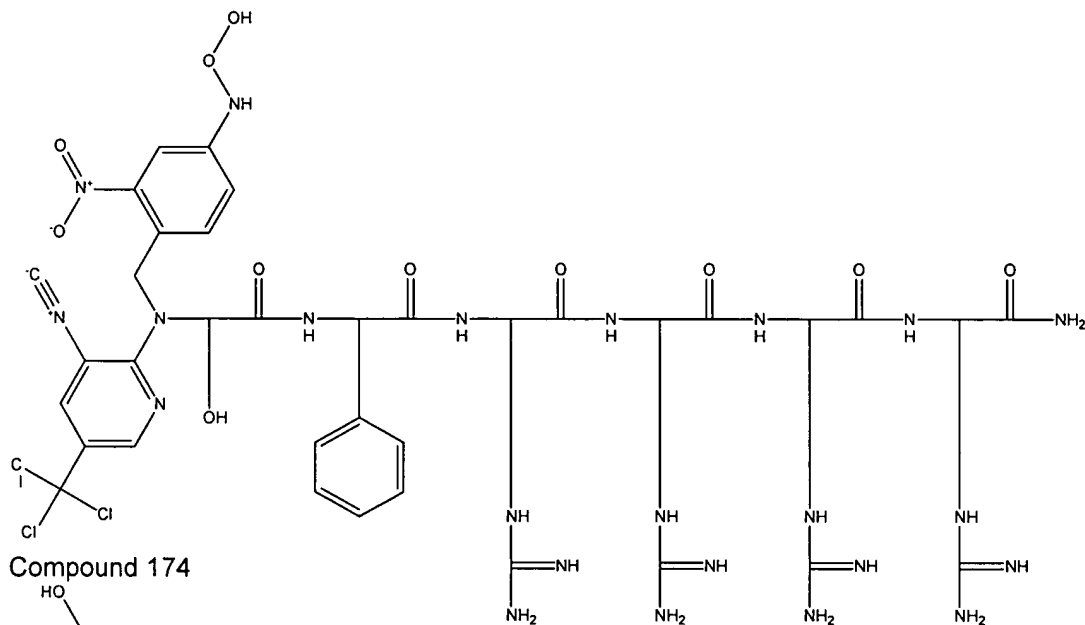
Compound 163

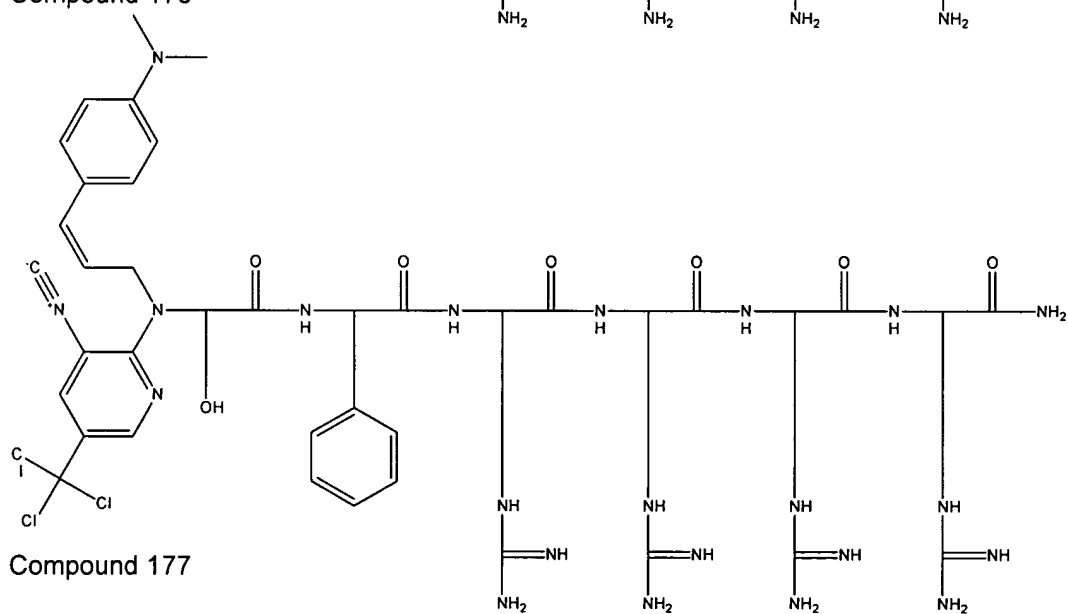
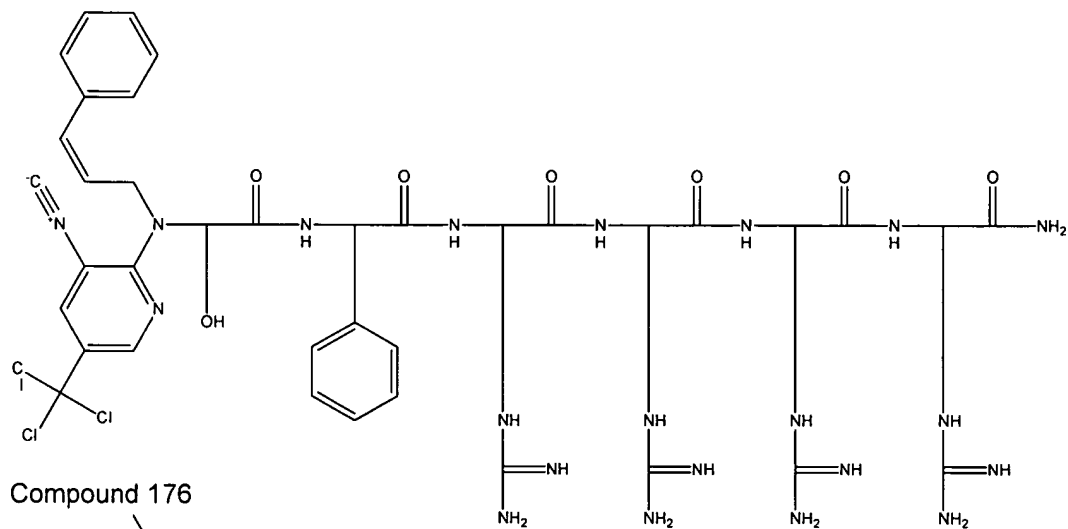




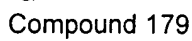
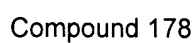


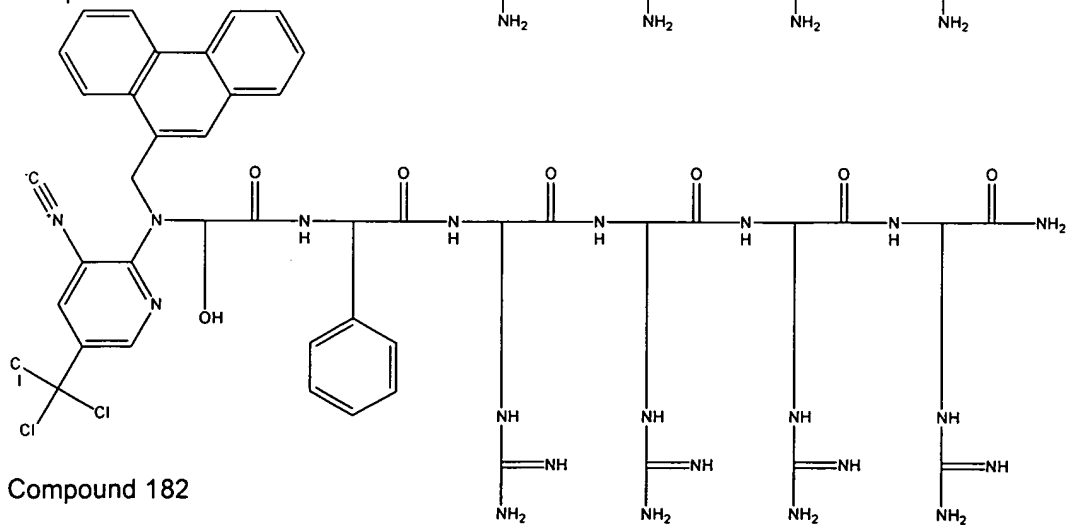
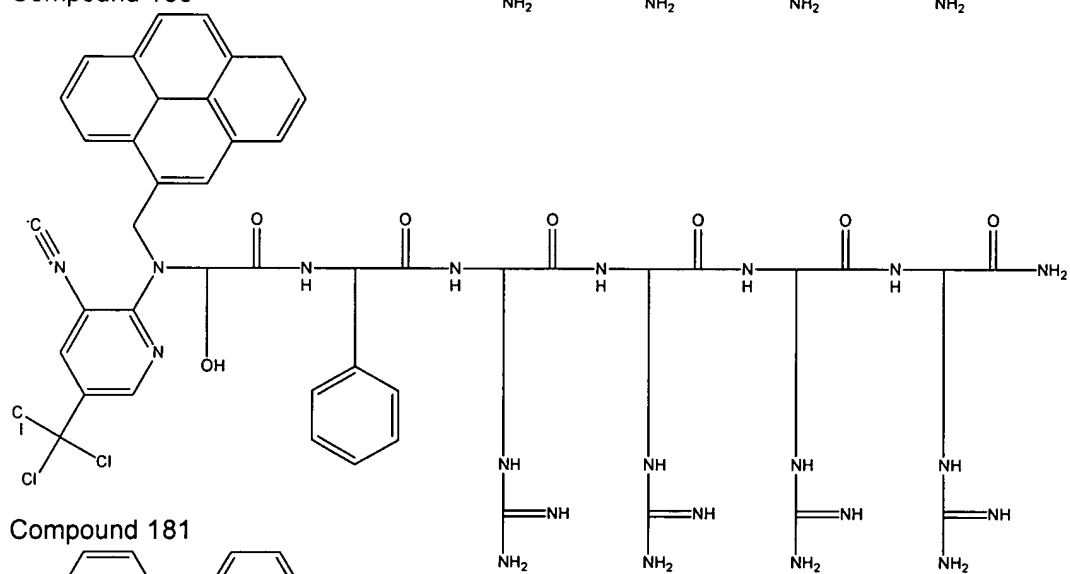
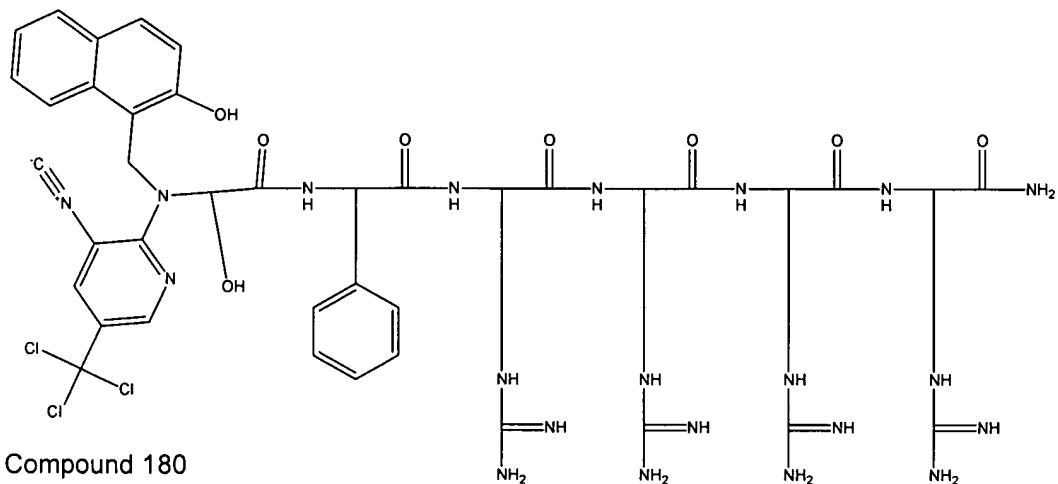


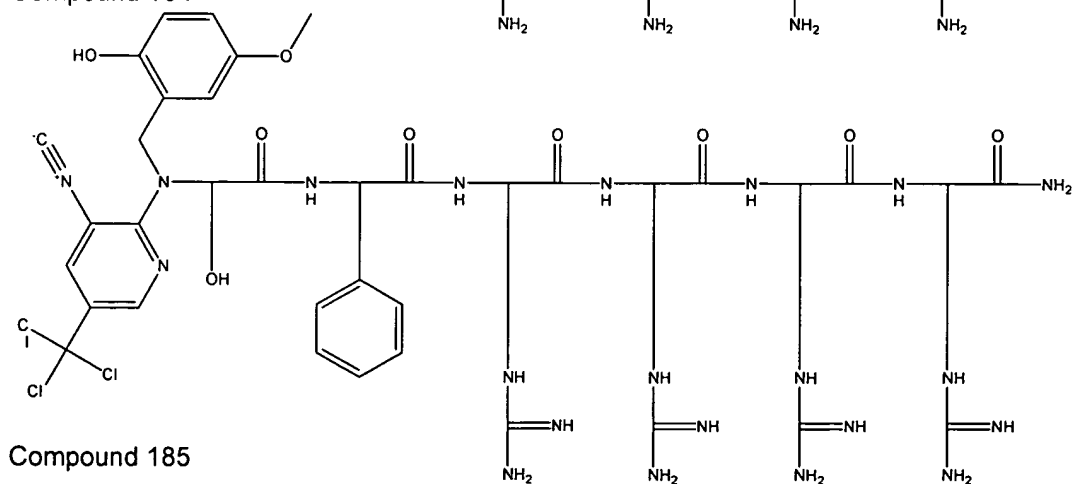
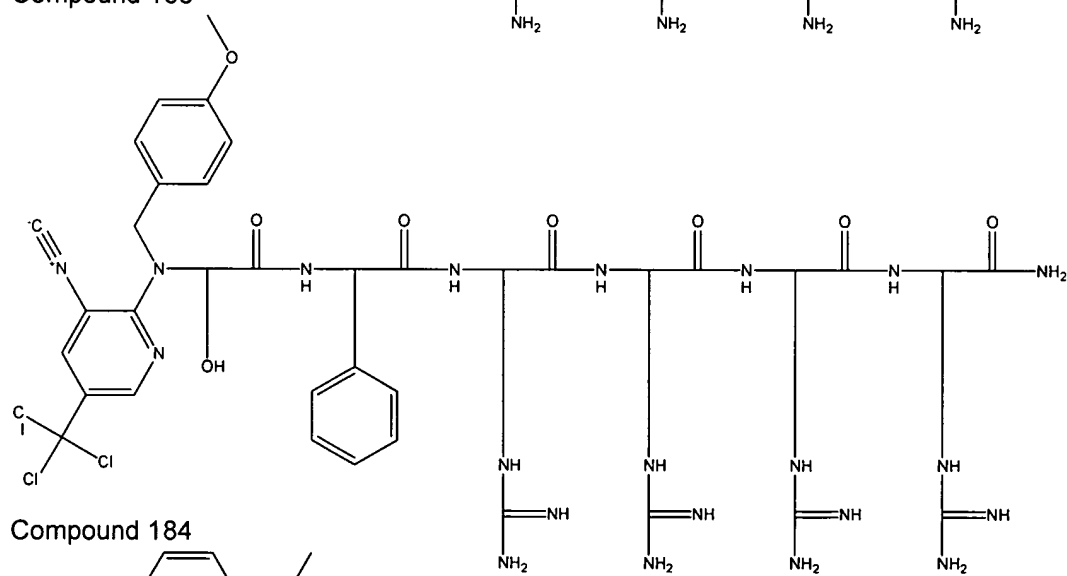
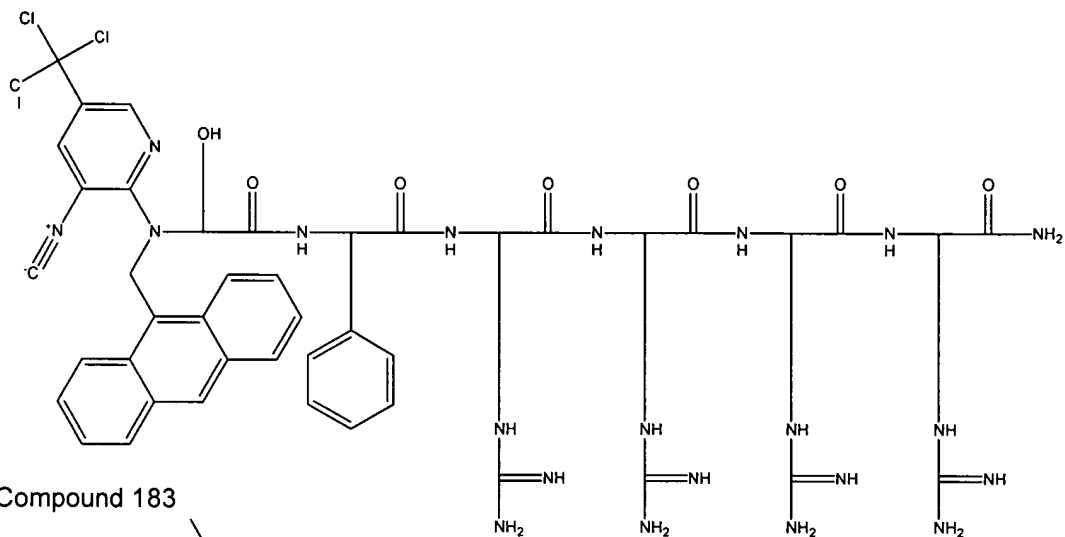




page 78 of 192





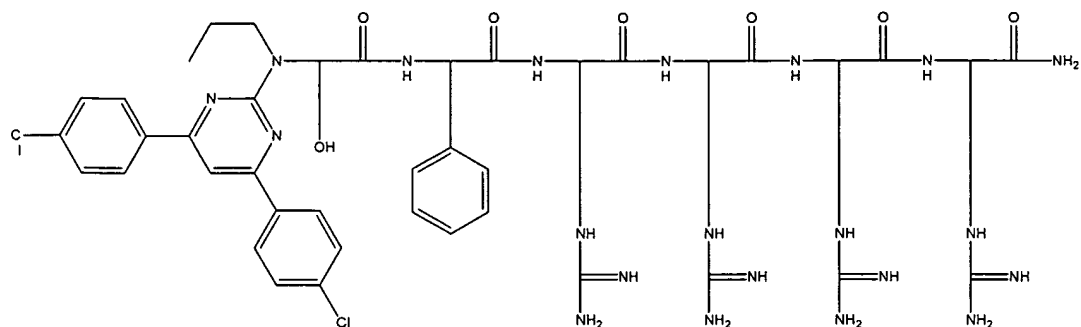


Applicant: David S. Lawrence

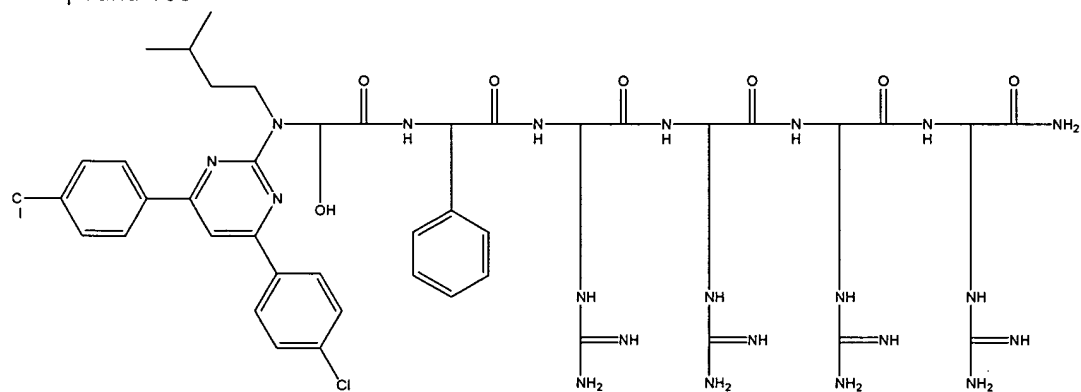
Serial No.: 10/755,086

Filed: January 9, 2004

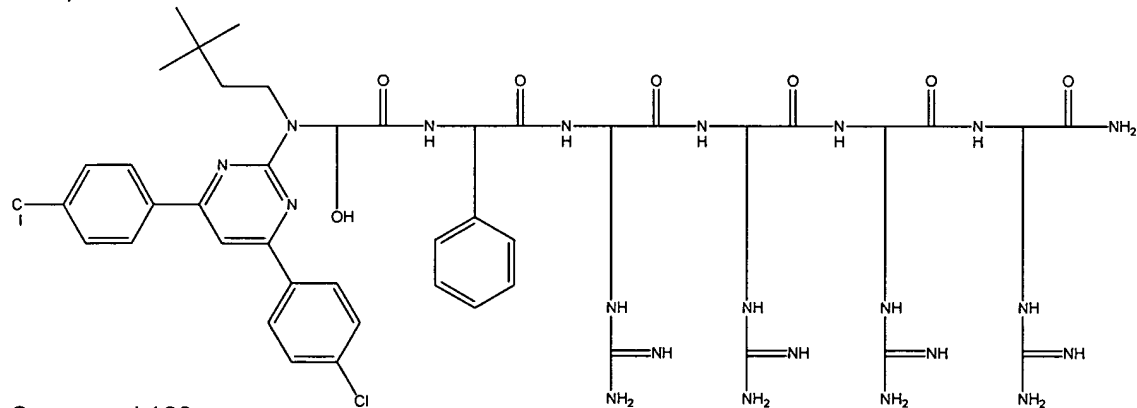
page 81 of 192



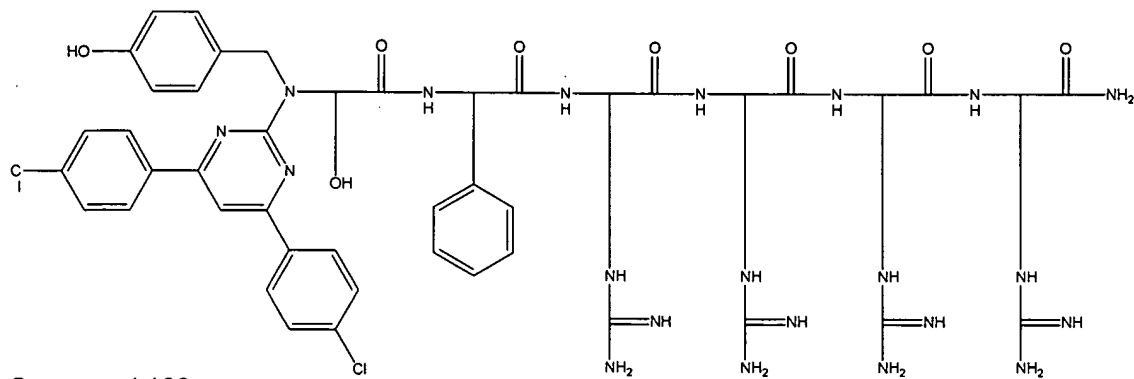
Compound 186



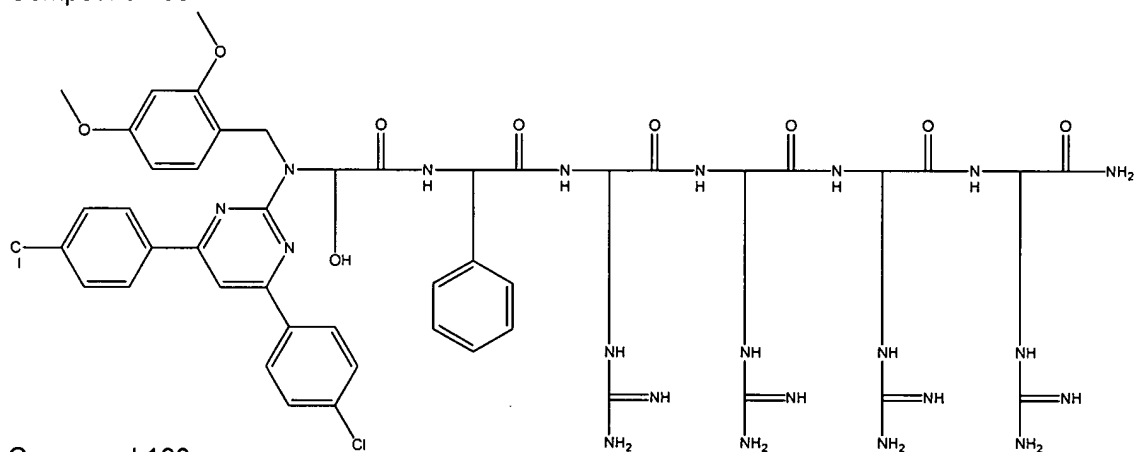
Compound 187



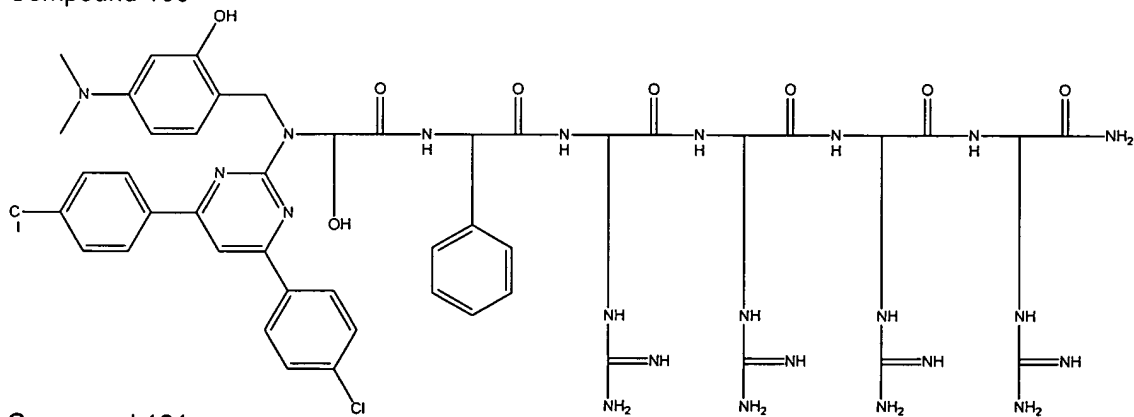
Compound 188



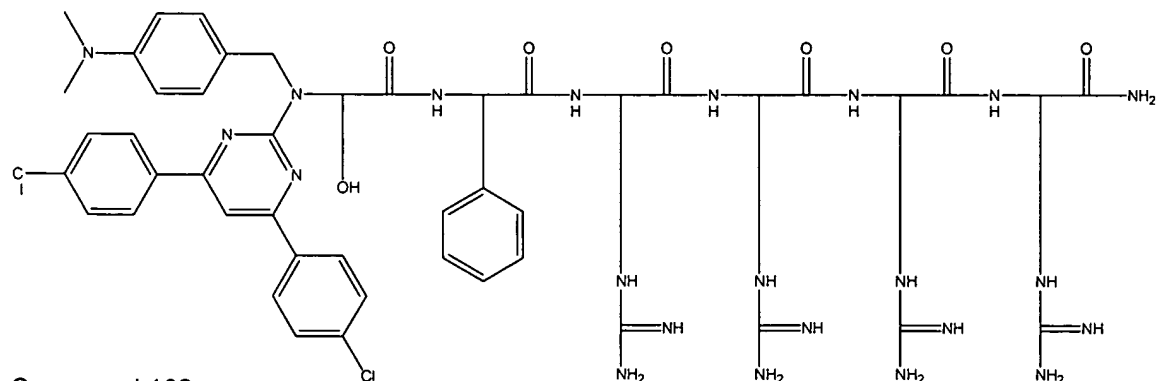
Compound 189



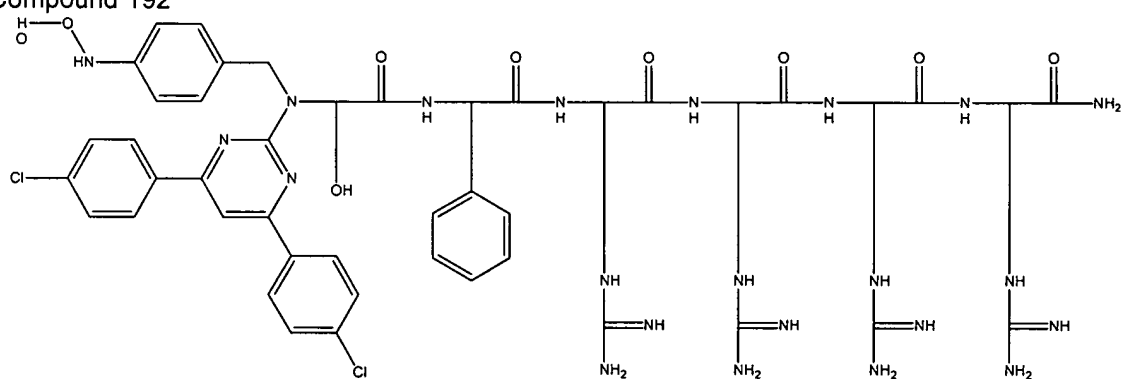
Compound 190



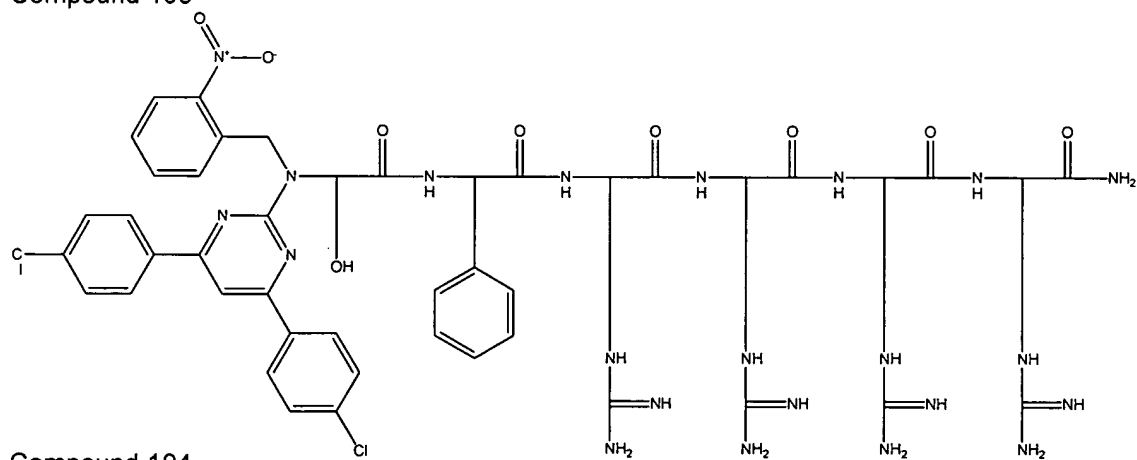
Compound 191



Compound 192



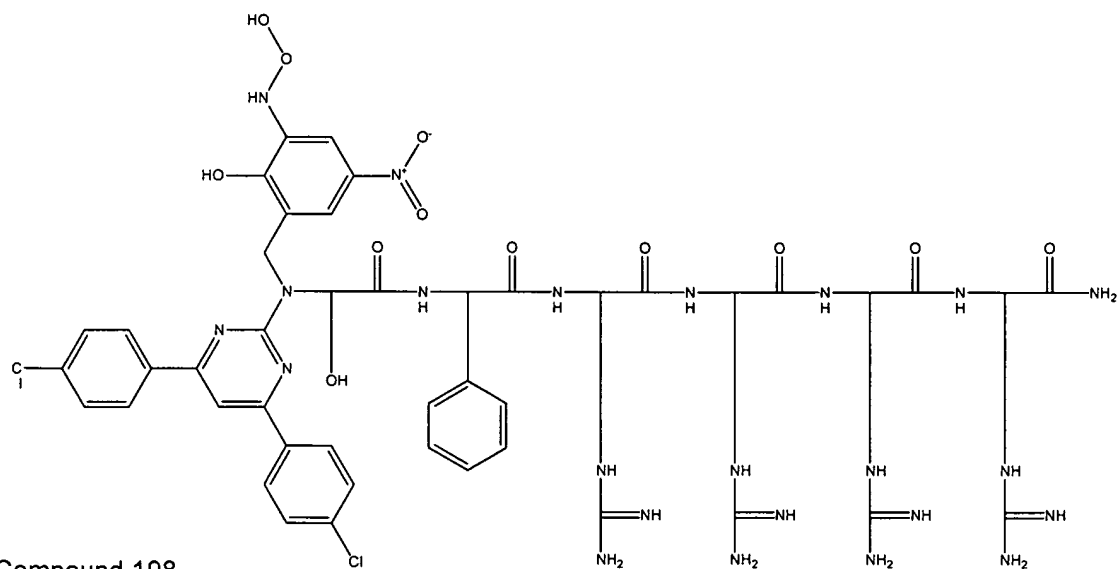
Compound 193



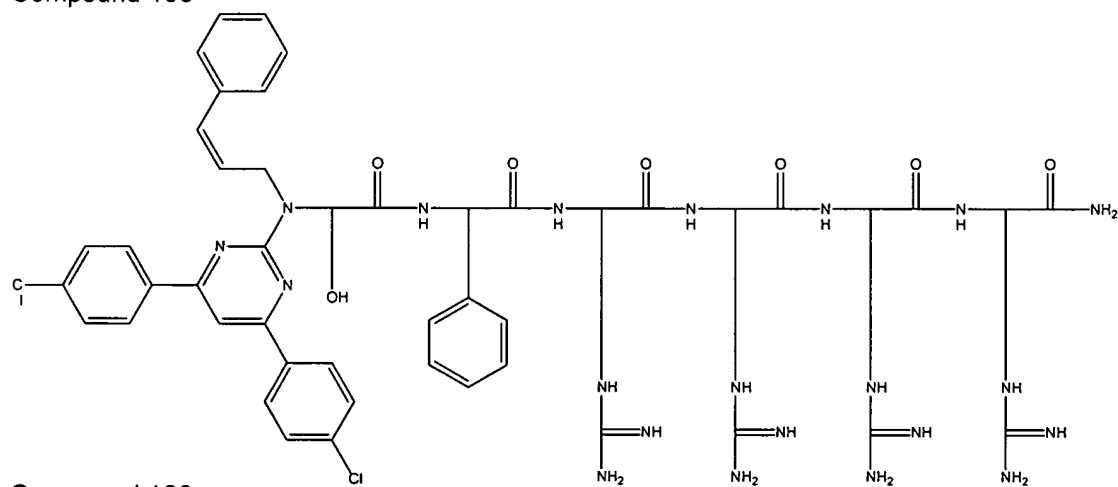
Compound 194

page 84 of 192



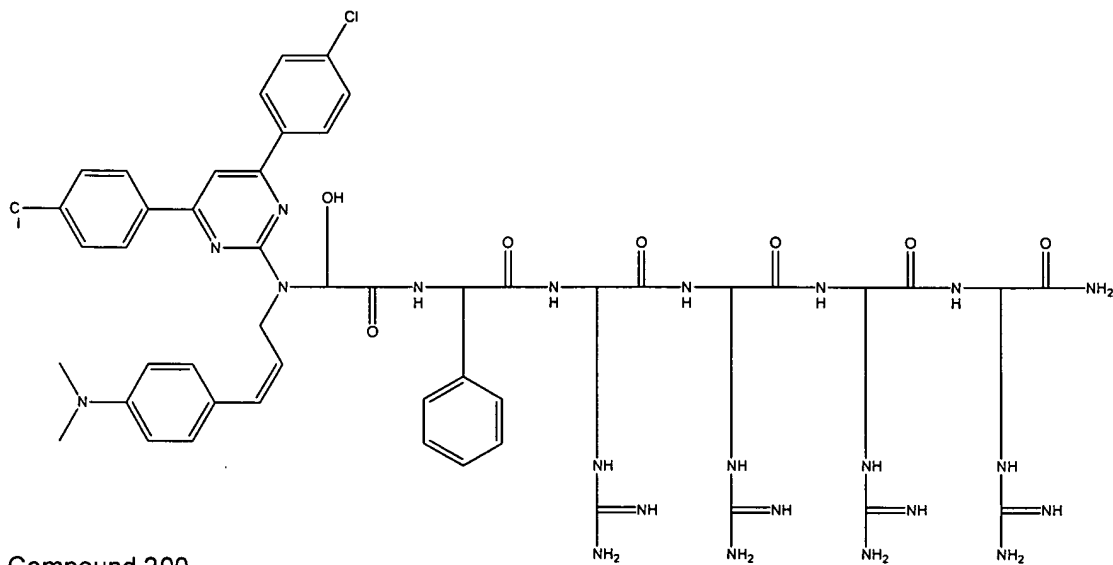


Compound 198

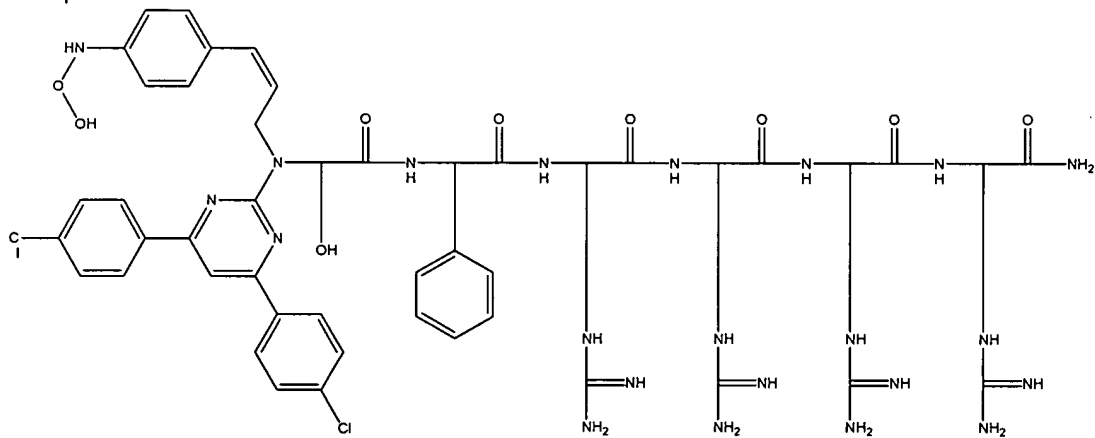


Compound 199

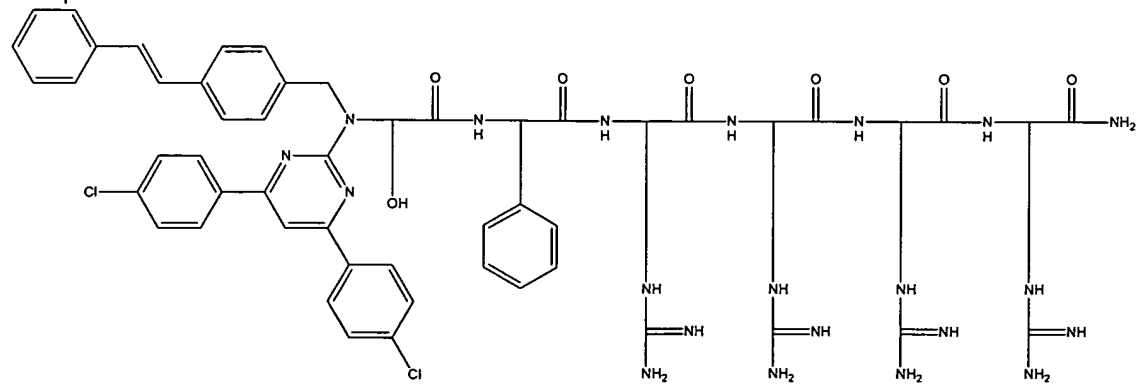
page 86 of 192



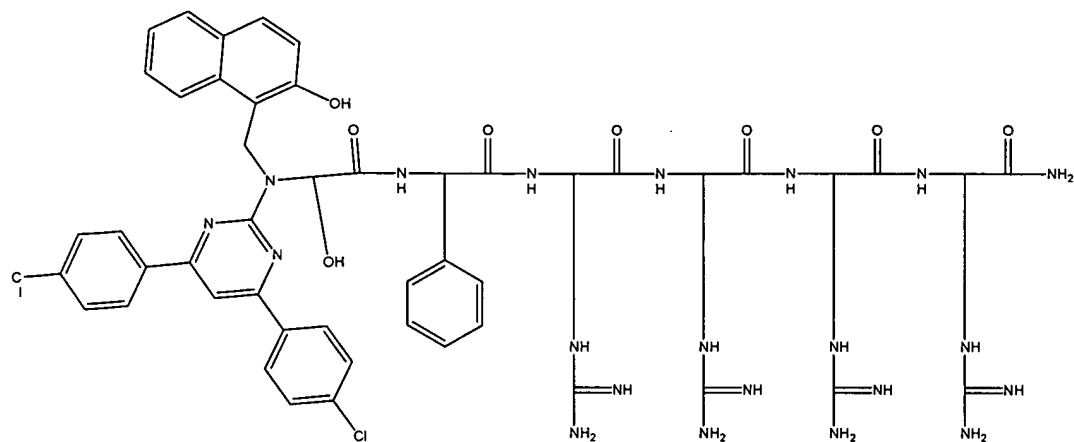
Compound 200



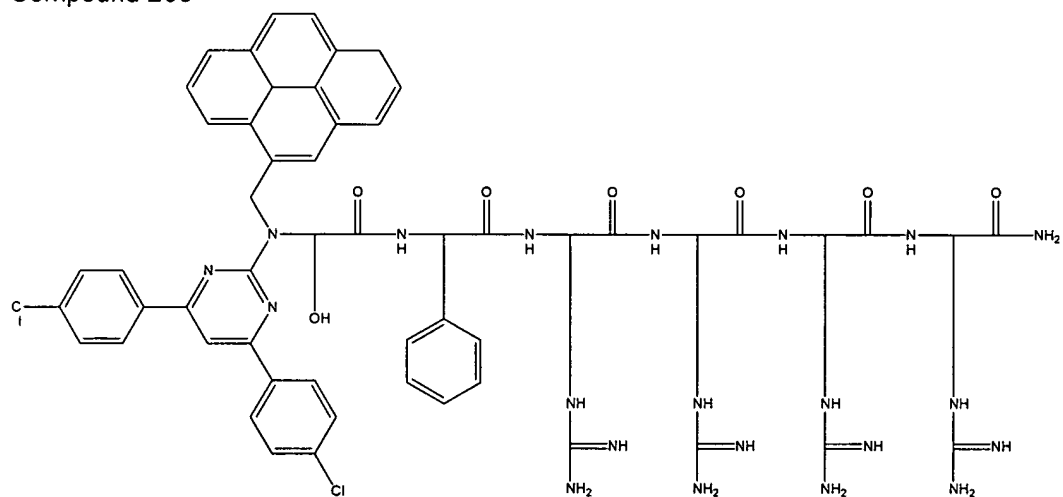
Compound 201



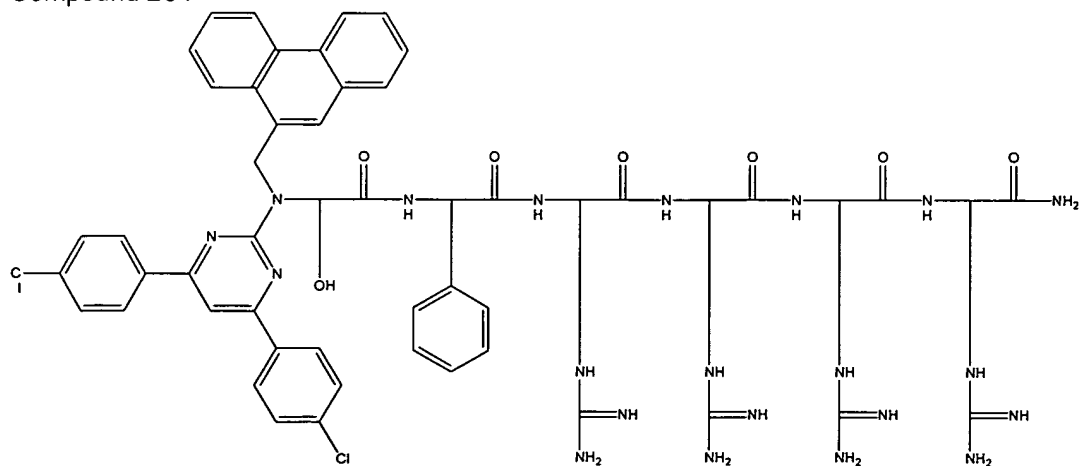
Compound 202



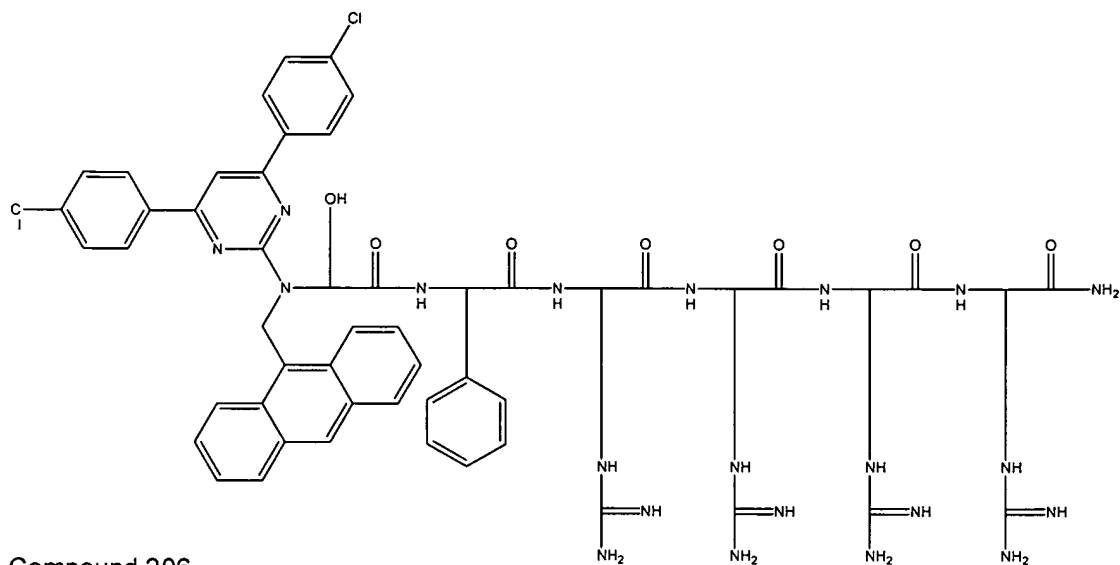
Compound 203



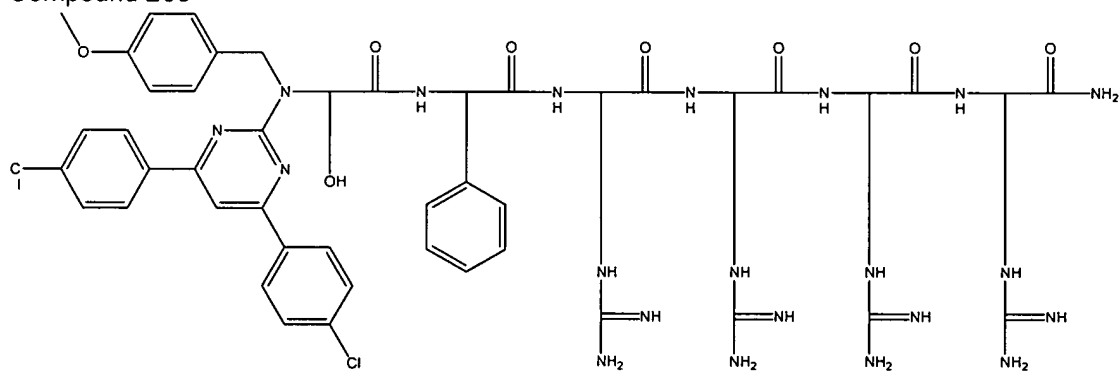
Compound 204



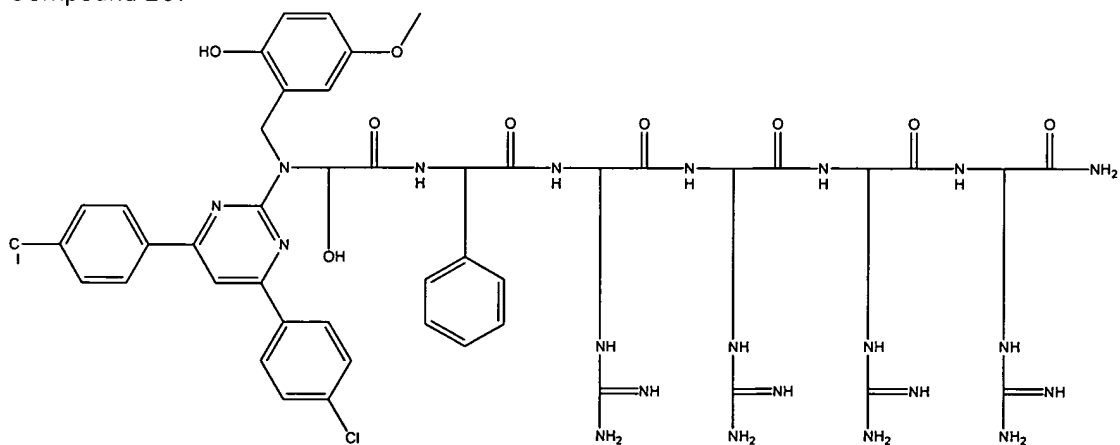
Compound 205



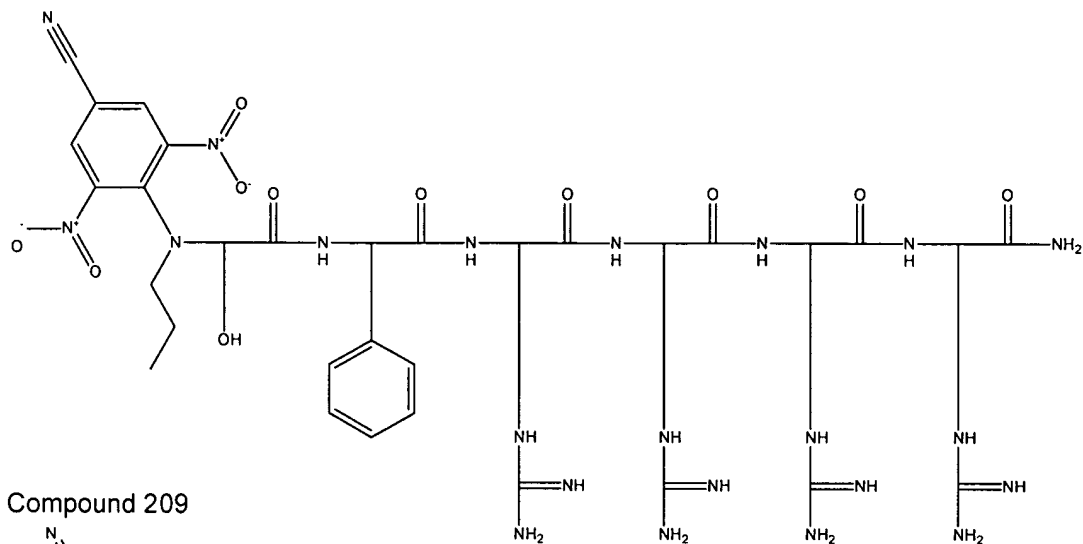
Compound 206



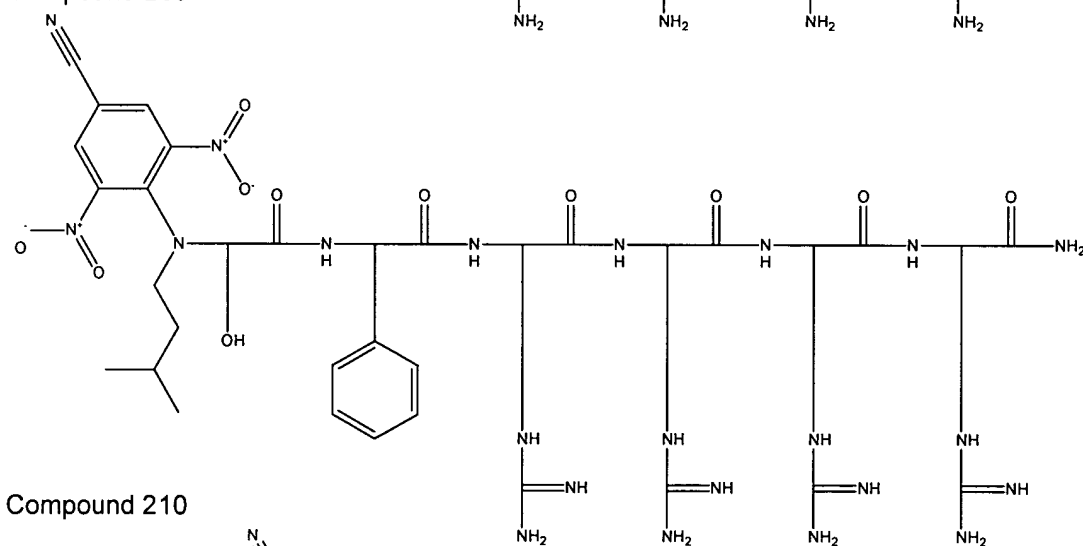
Compound 207



Compound 208

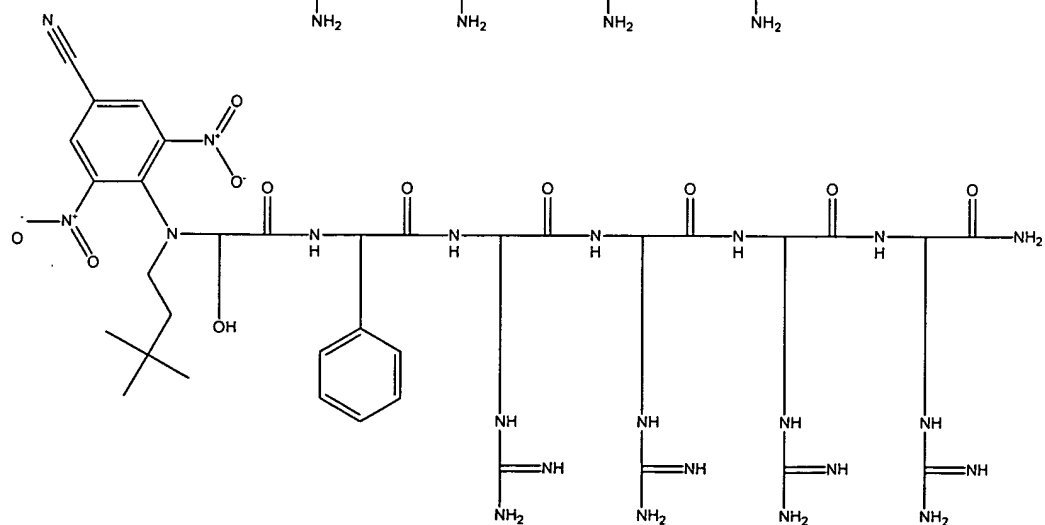


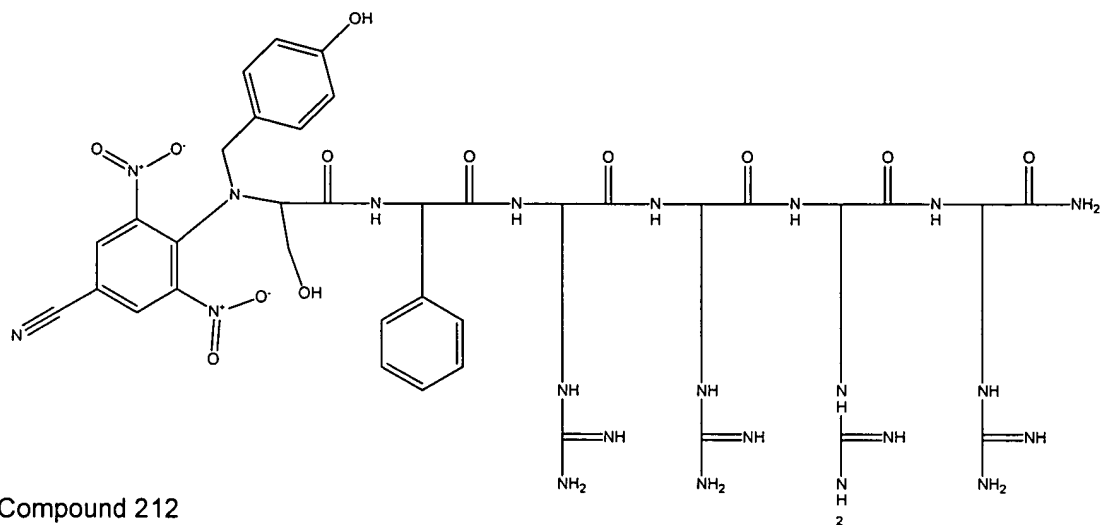
Compound 209



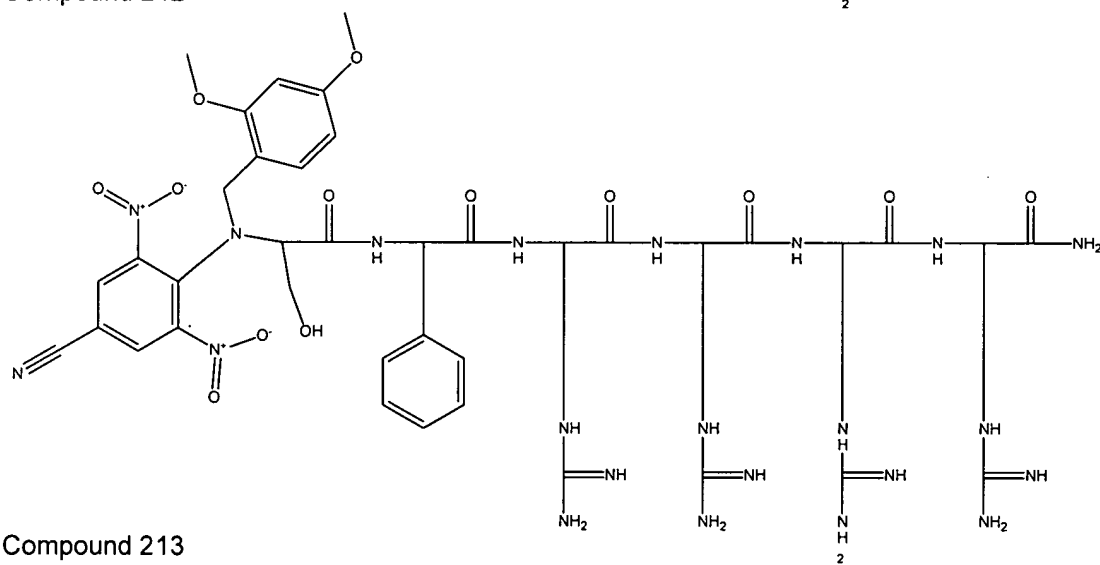
Compound 210

Compound 211





Compound 212



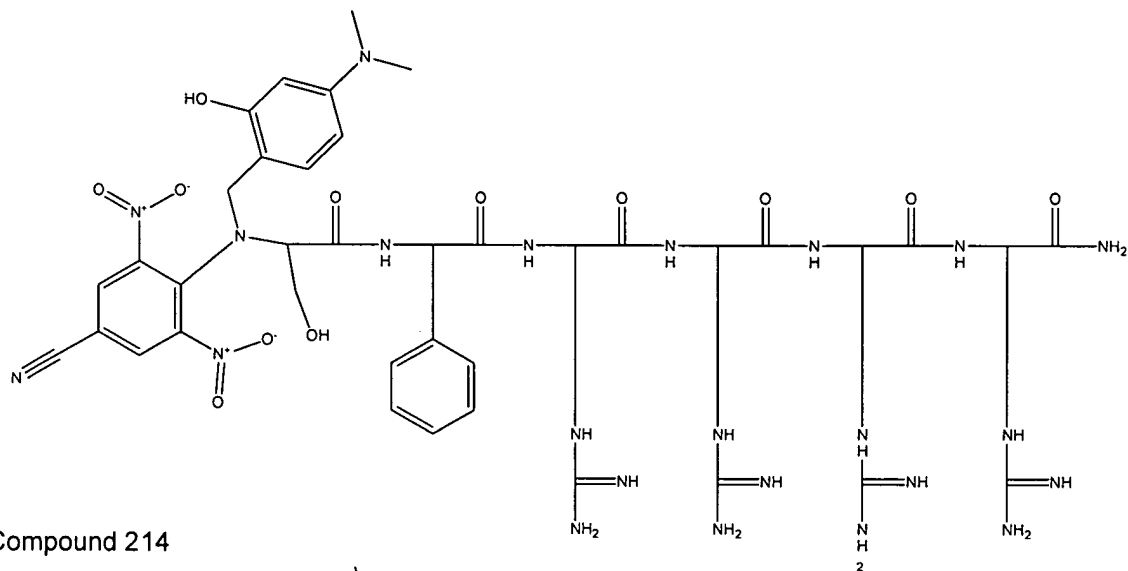
Compound 213

Applicant: David S. Lawrence

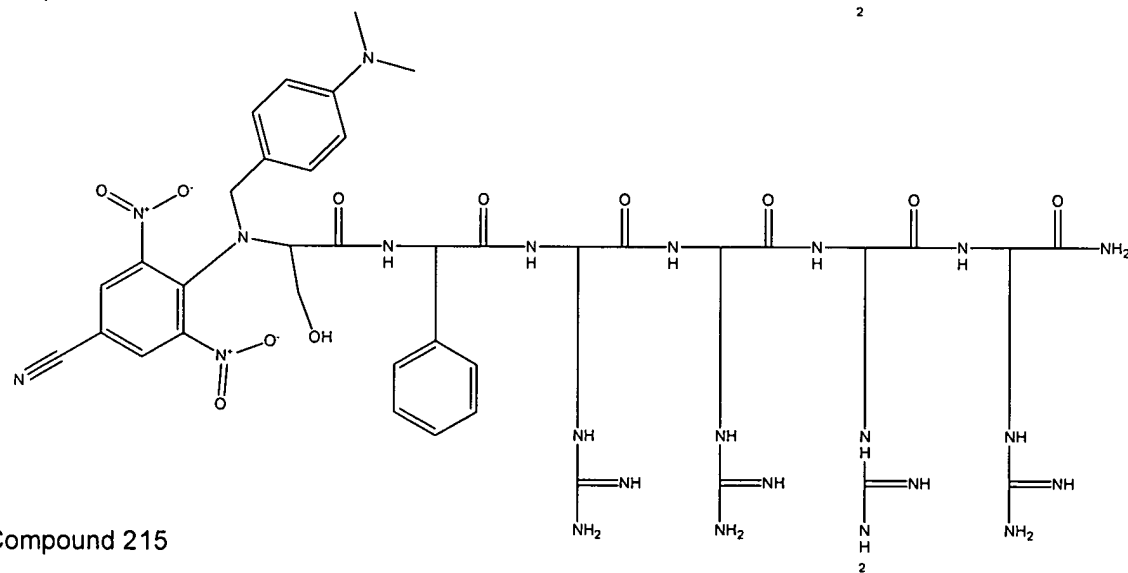
Serial No.: 10/755,086

Filed: January 9, 2004

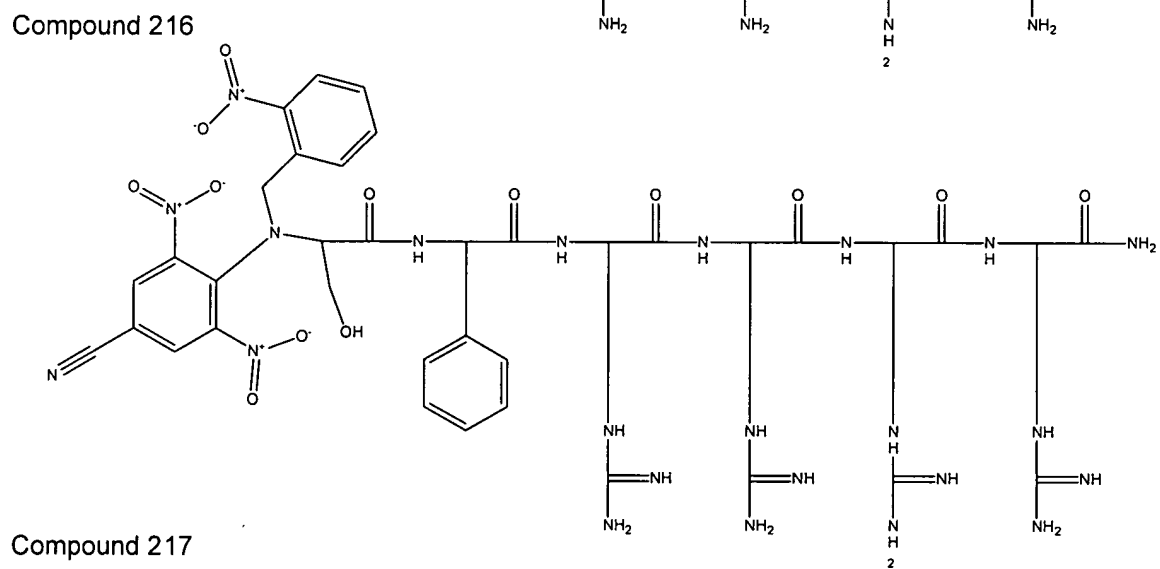
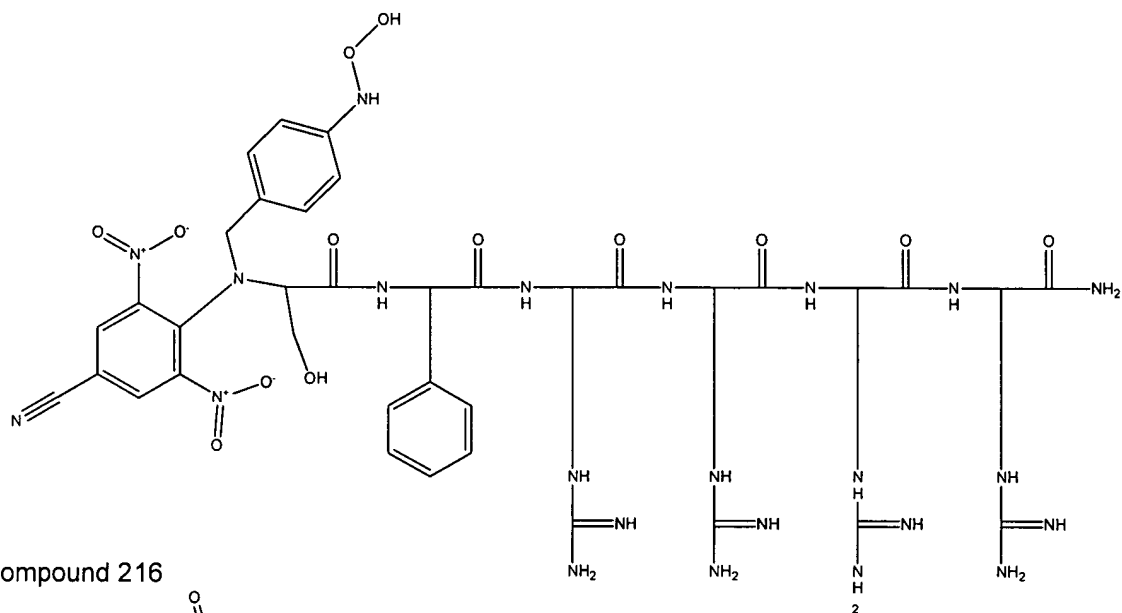
page 91 of 192



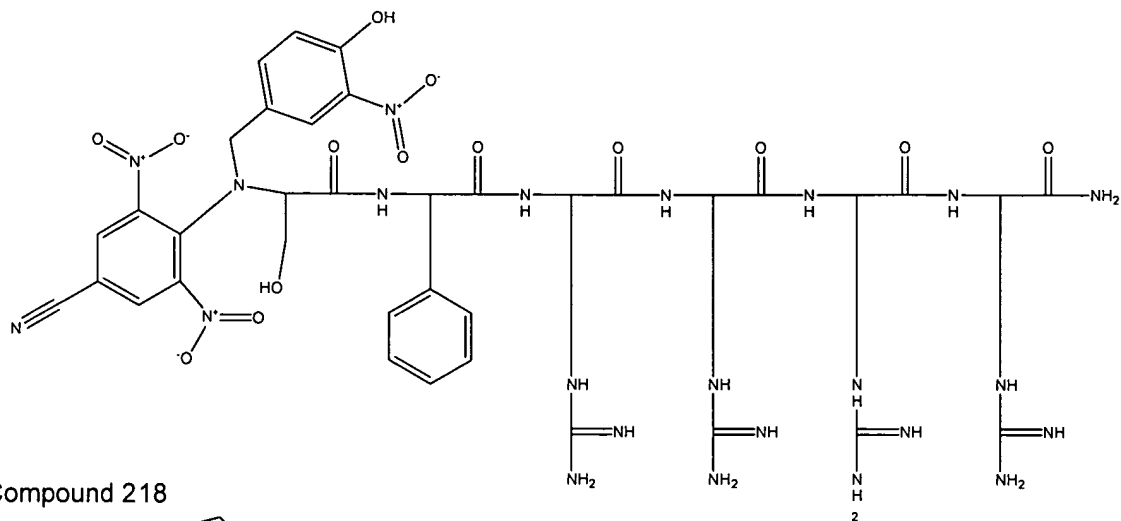
Compound 214



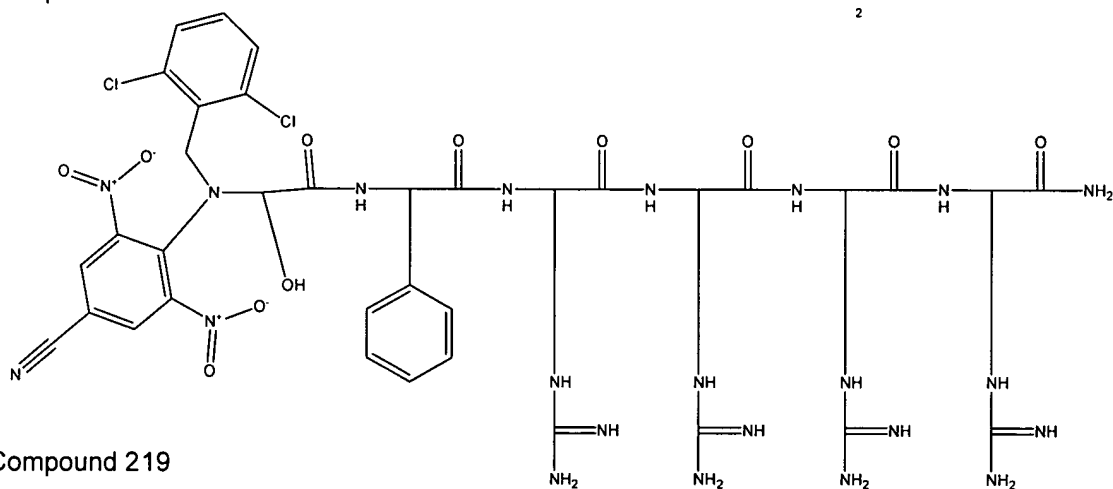
Compound 215



Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 93 of 192

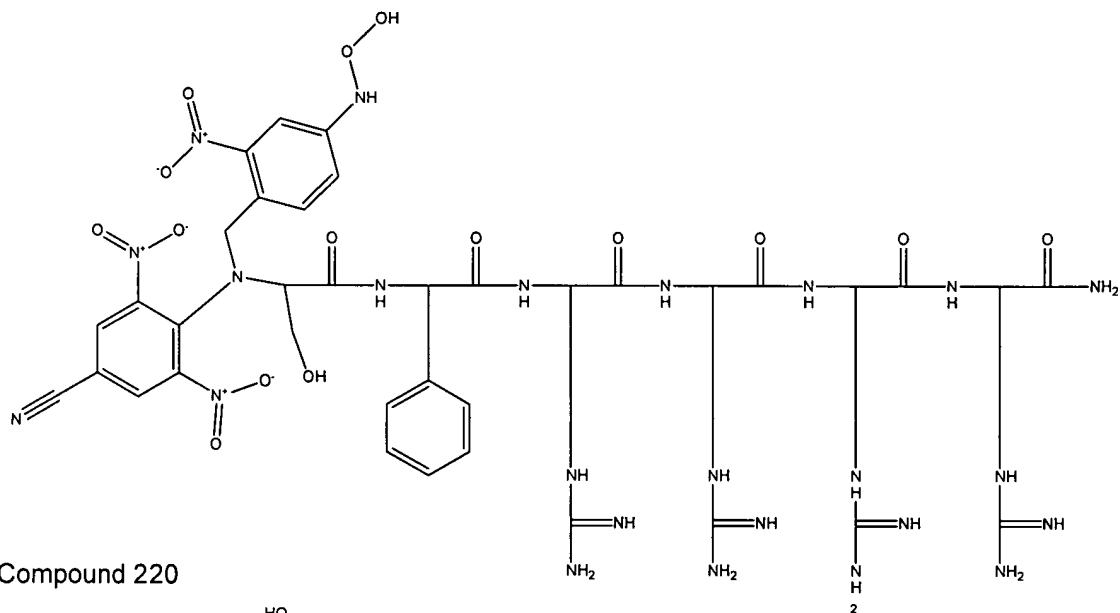


Compound 218

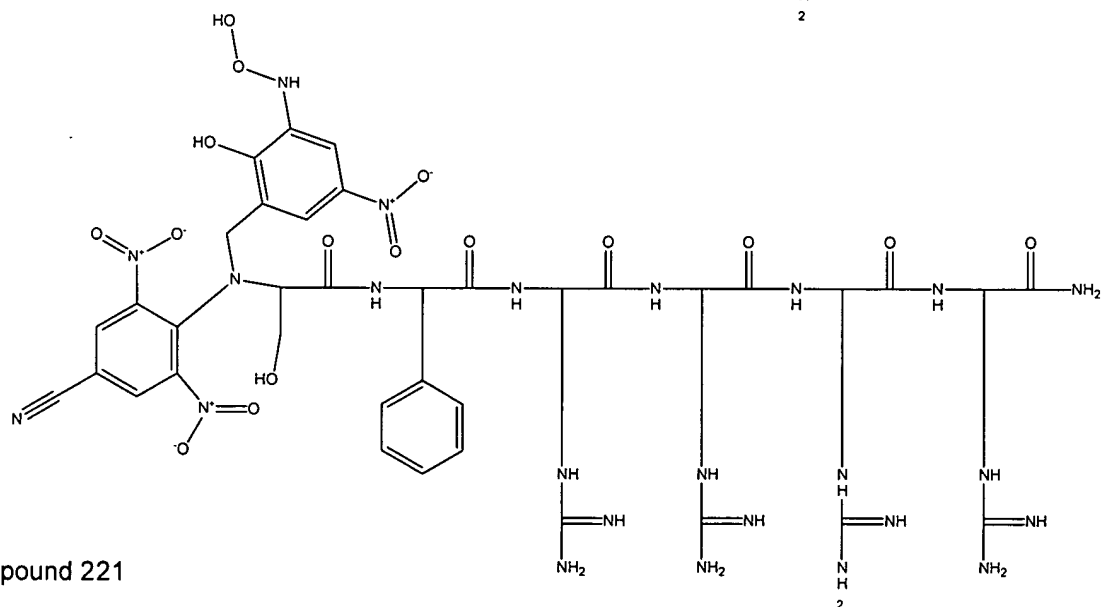


Compound 219

397236.1



Compound 220



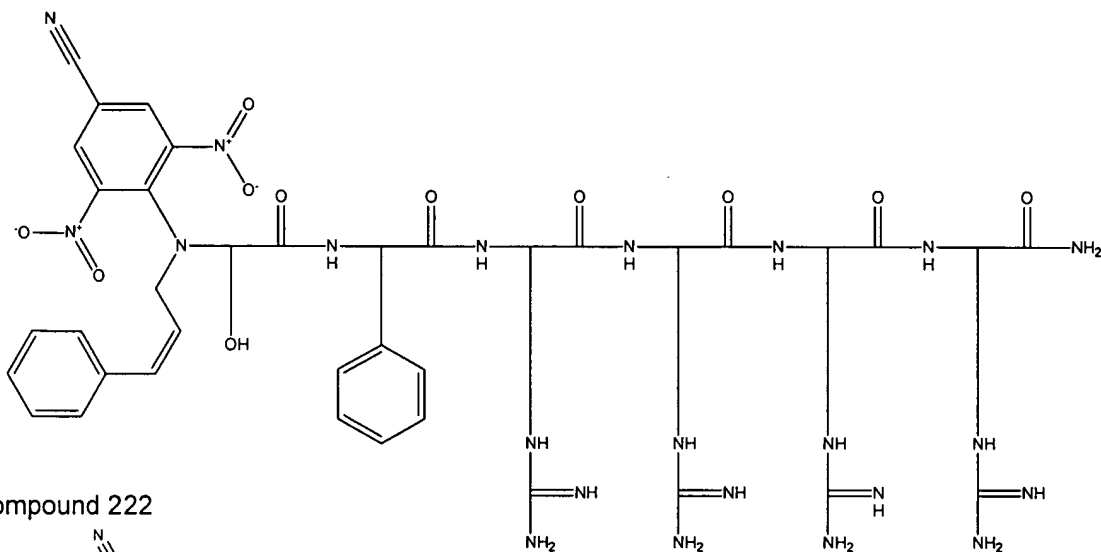
Compound 221

Applicant: David S. Lawrence

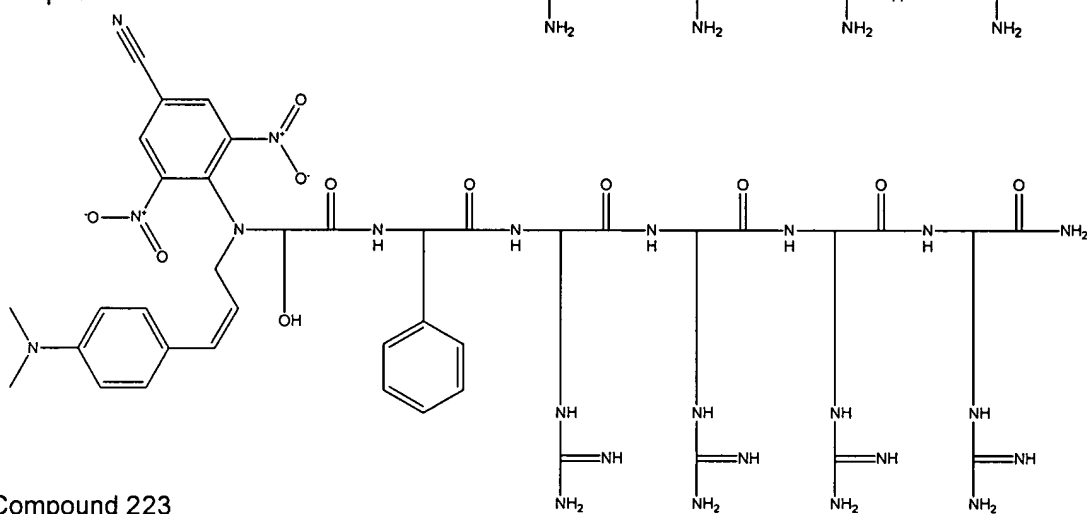
Serial No.: 10/755,086

Filed: January 9, 2004

page 95 of 192

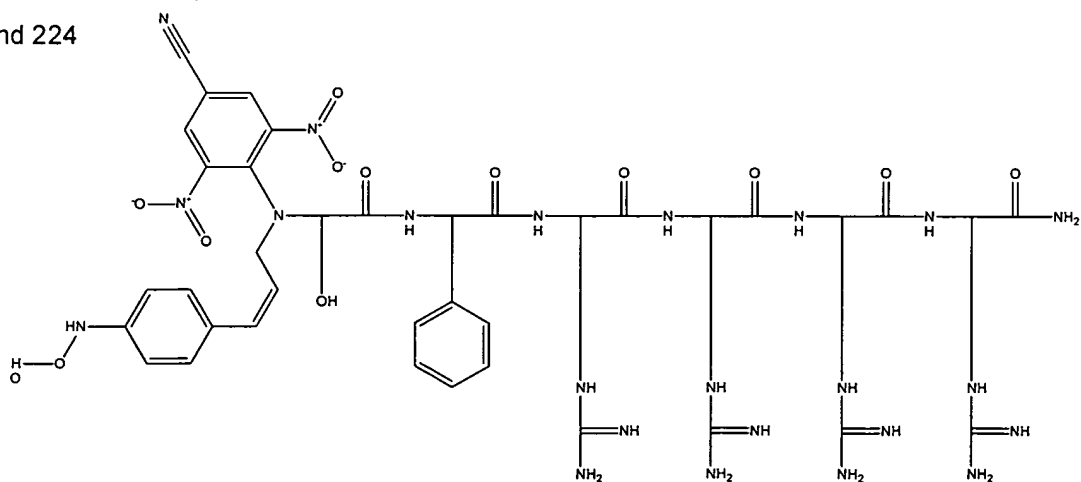


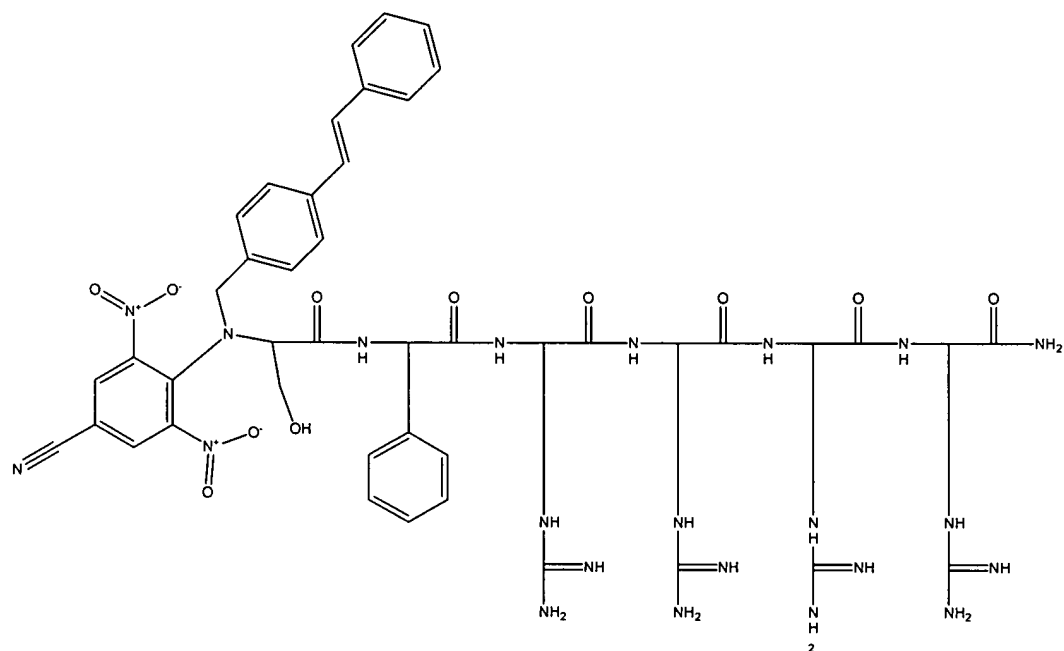
Compound 222



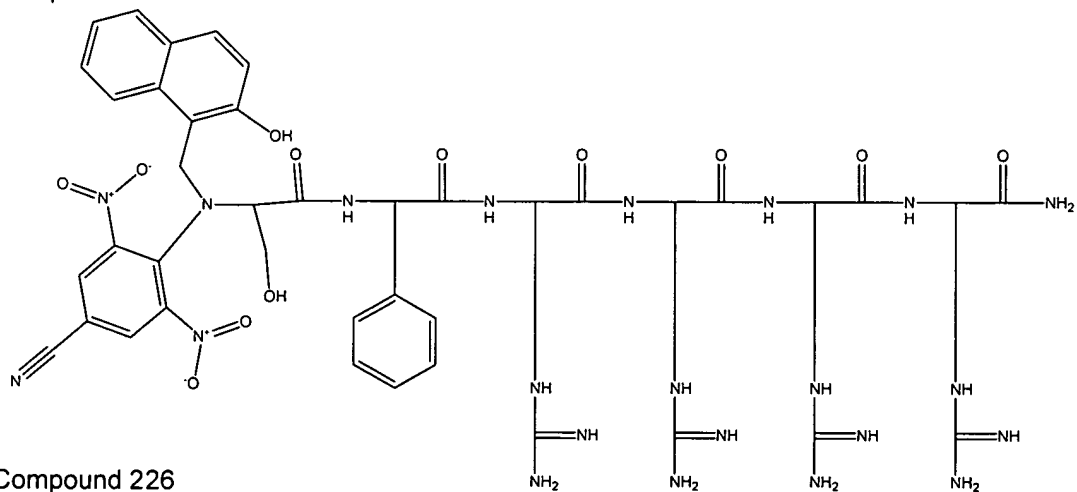
Compound 223

Compound 224



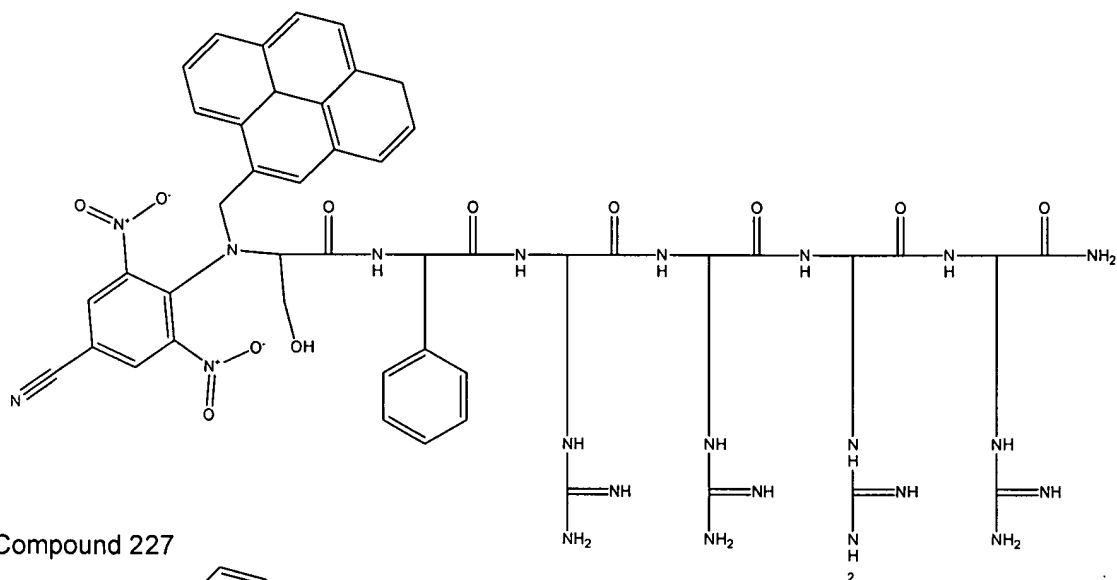


Compound 225

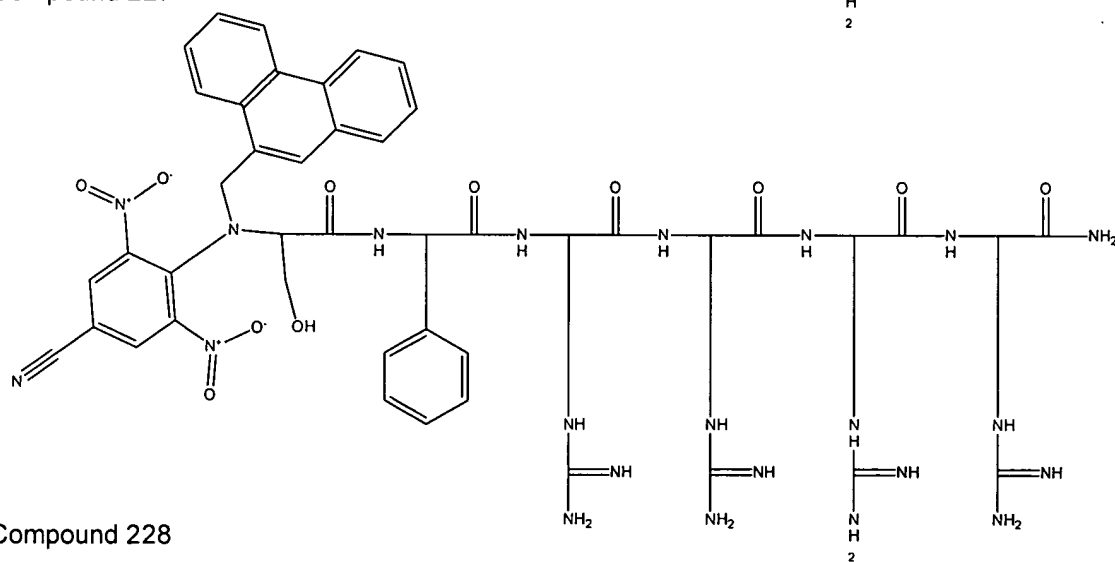


Compound 226

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 97 of 192

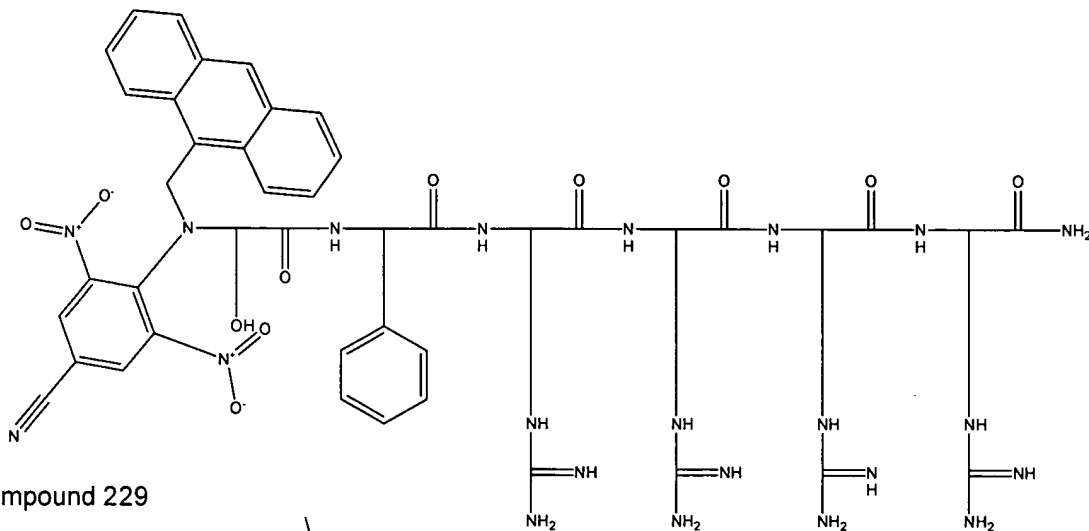


Compound 227

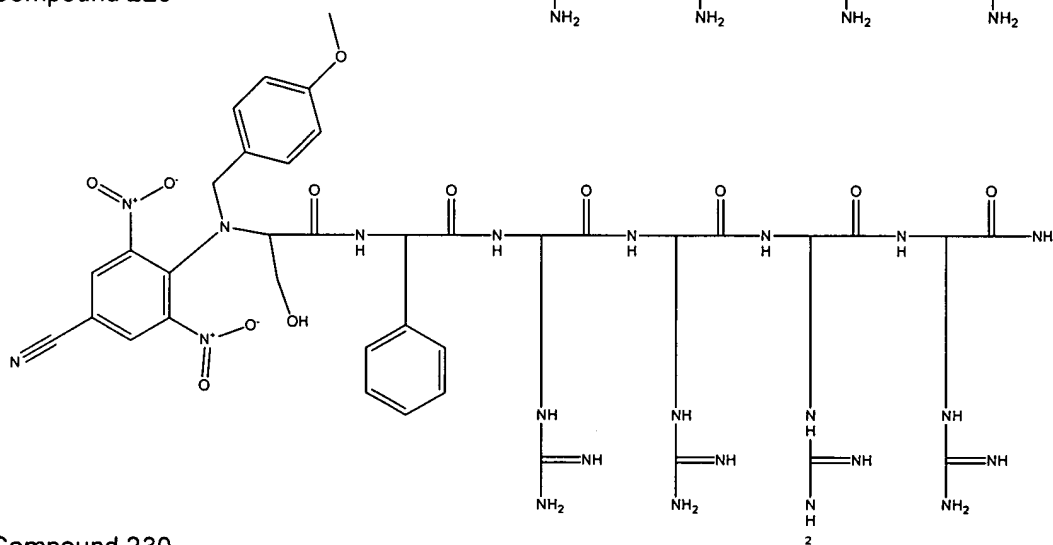


Compound 228

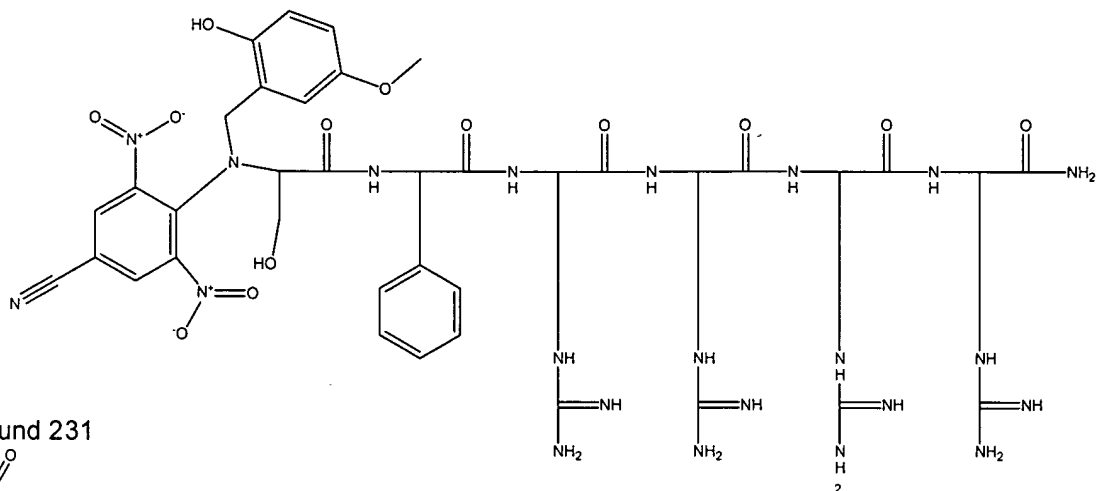
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 98 of 192



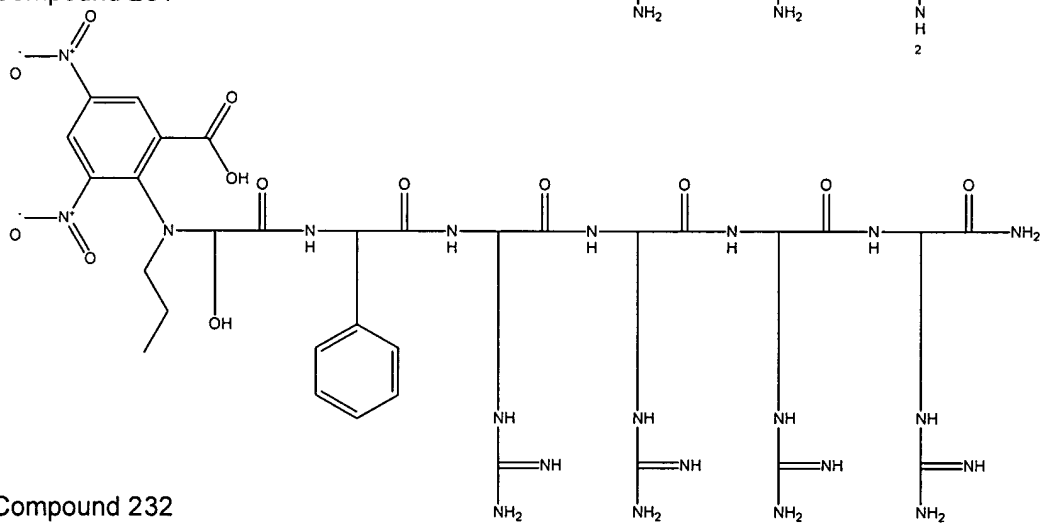
Compound 229



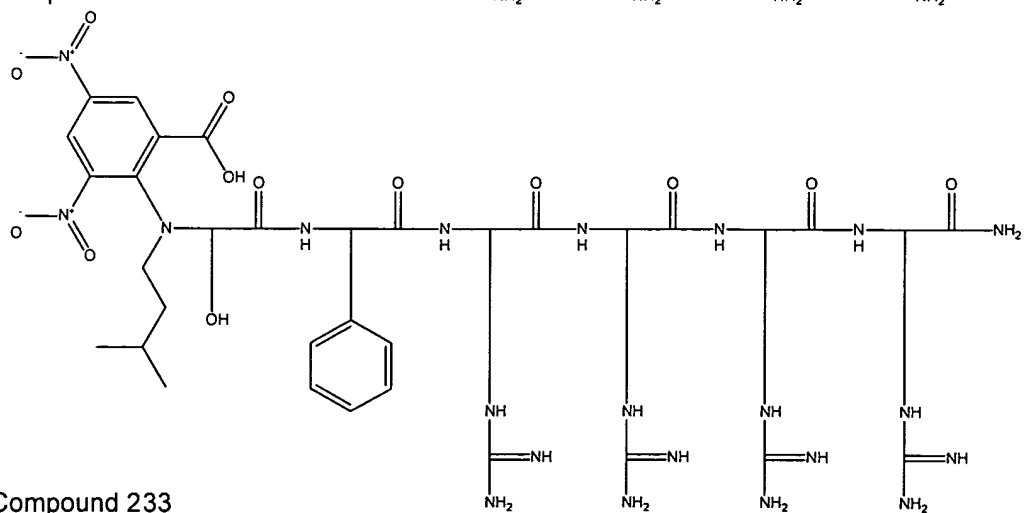
Compound 230



Compound 231

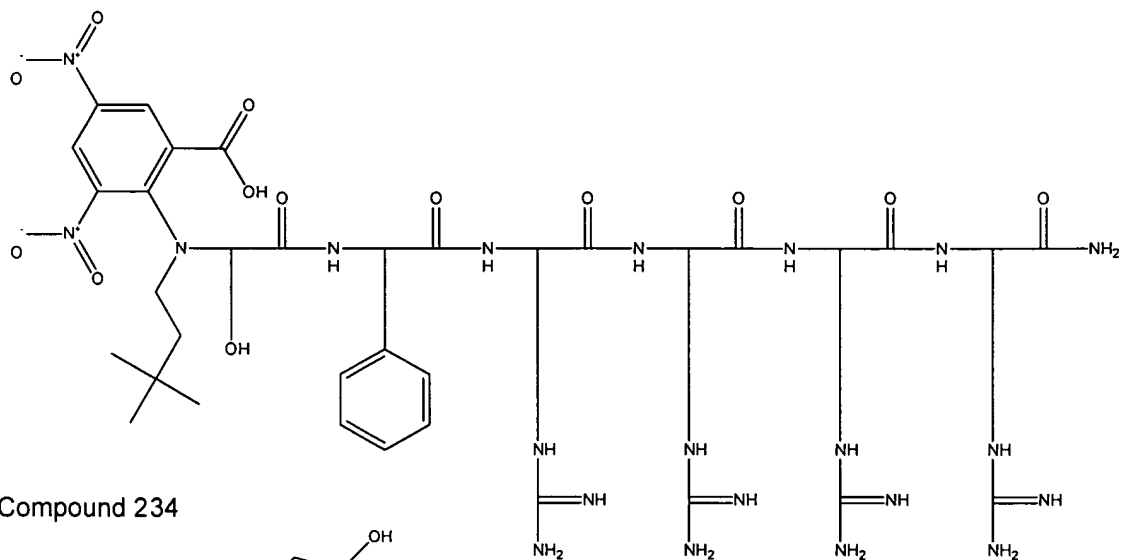


Compound 232

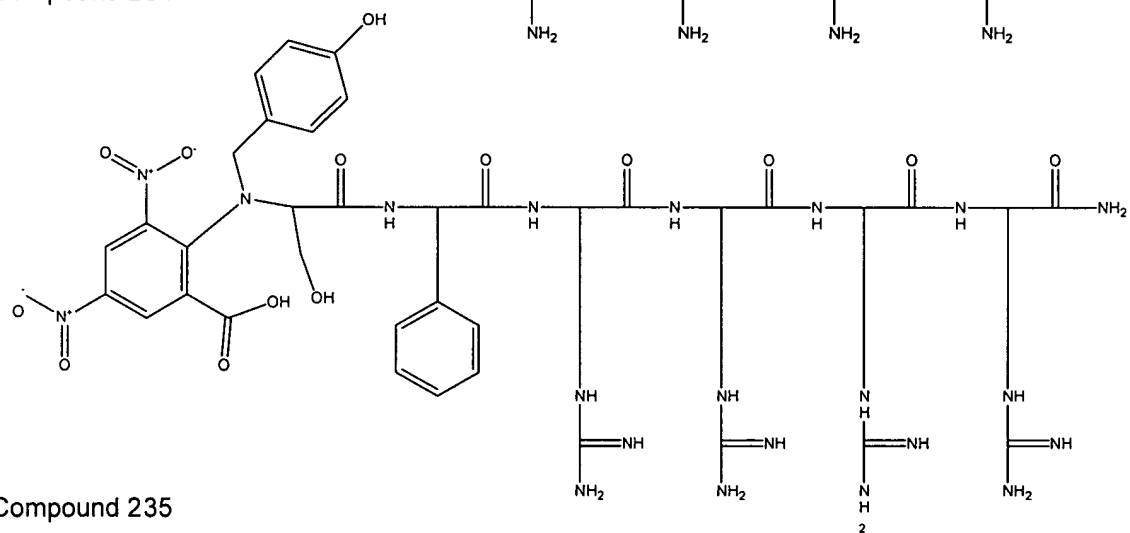


Compound 233

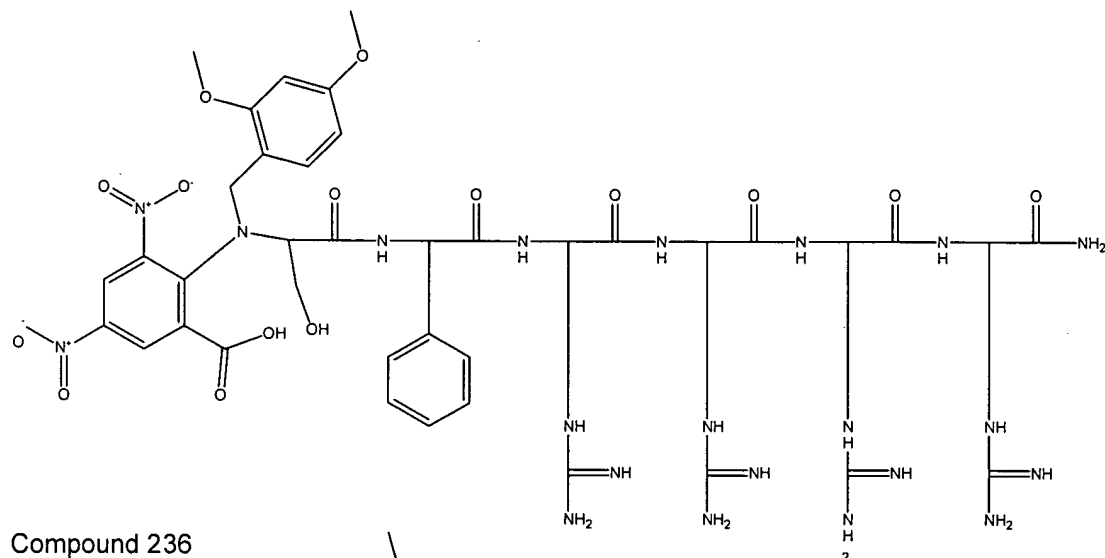
Applicant: David S. Lawrence
 Serial No.: 10/755,086
 Filed: January 9, 2004
 page 100 of 192



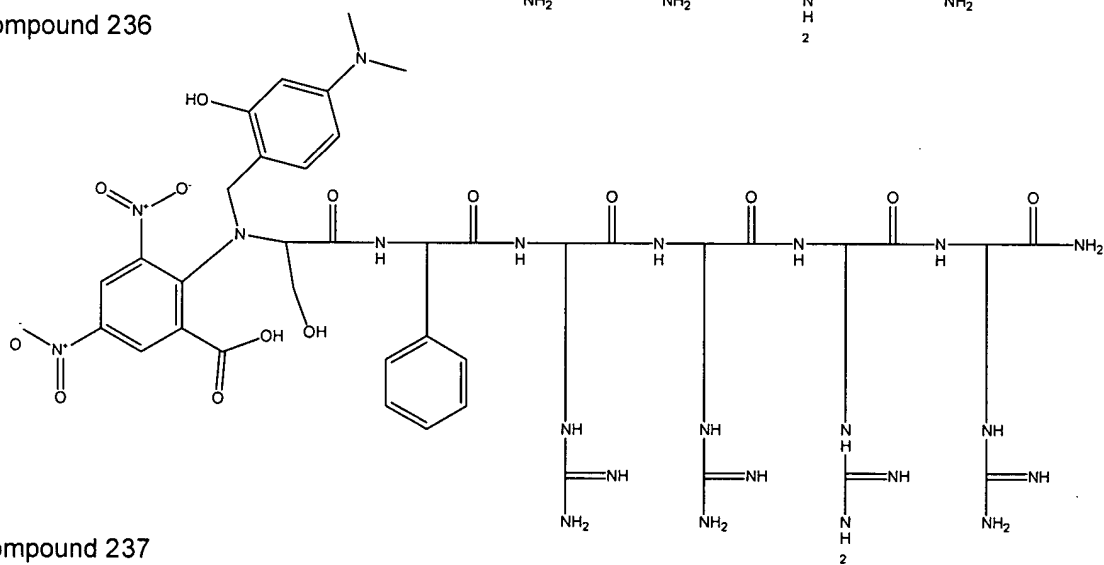
Compound 234



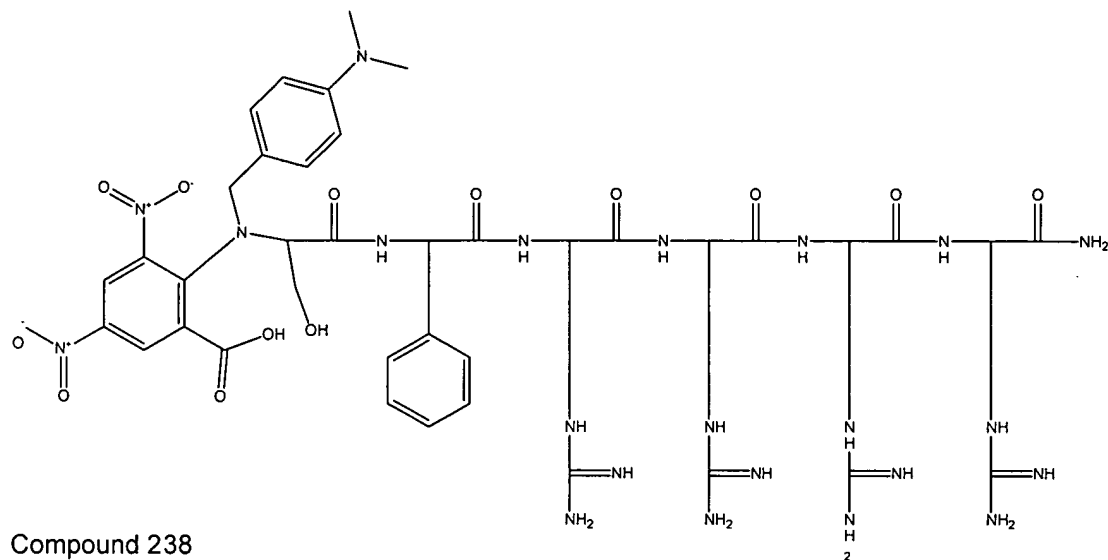
Compound 235



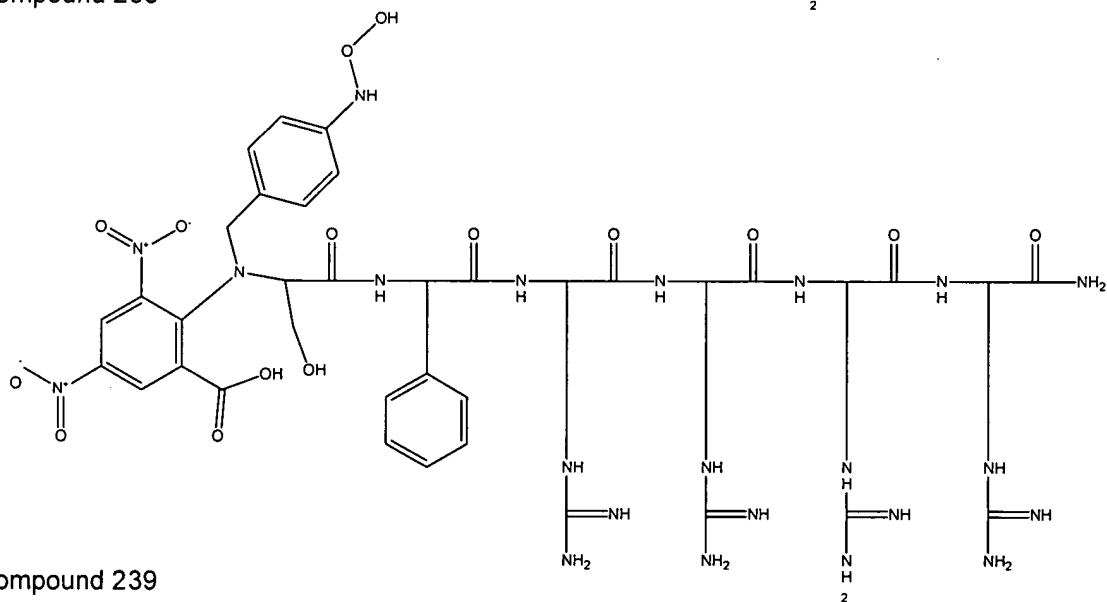
Compound 236



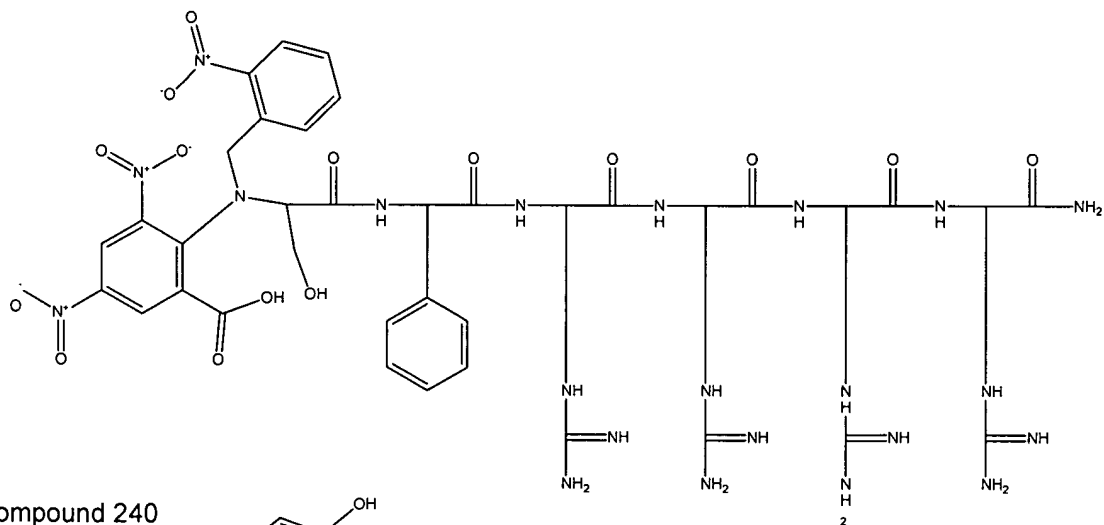
Compound 237



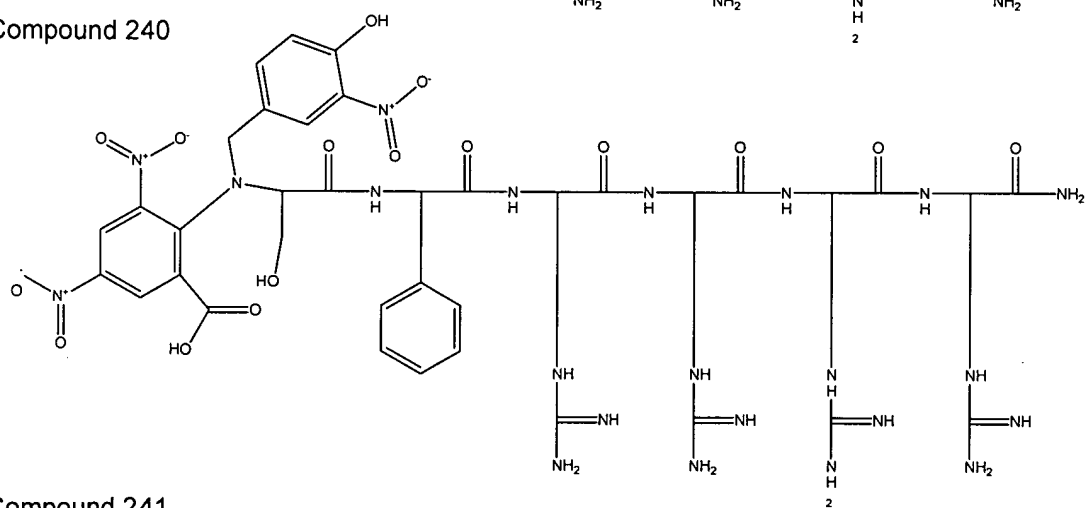
Compound 238



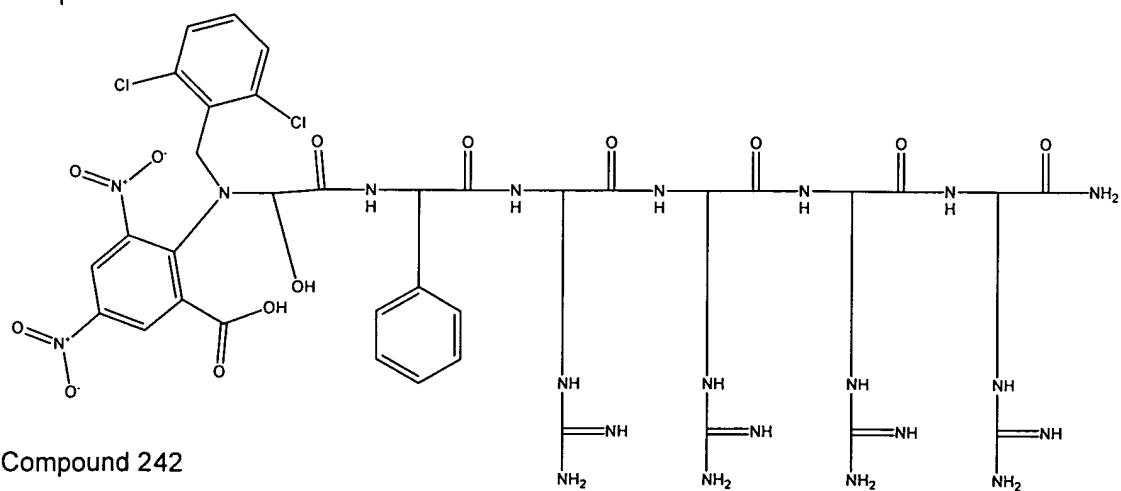
Compound 239



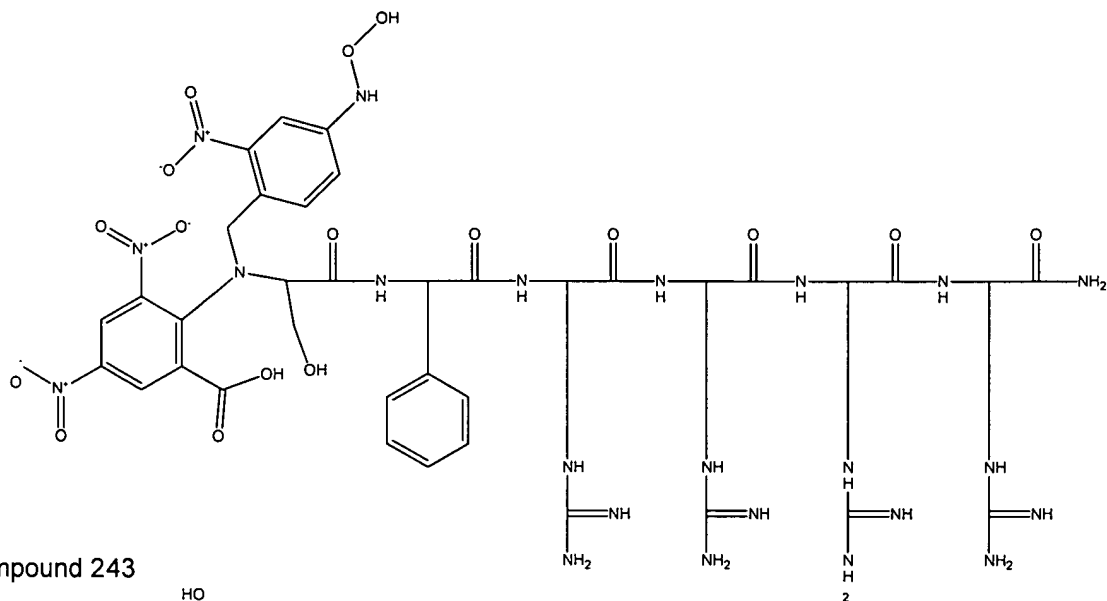
Compound 240



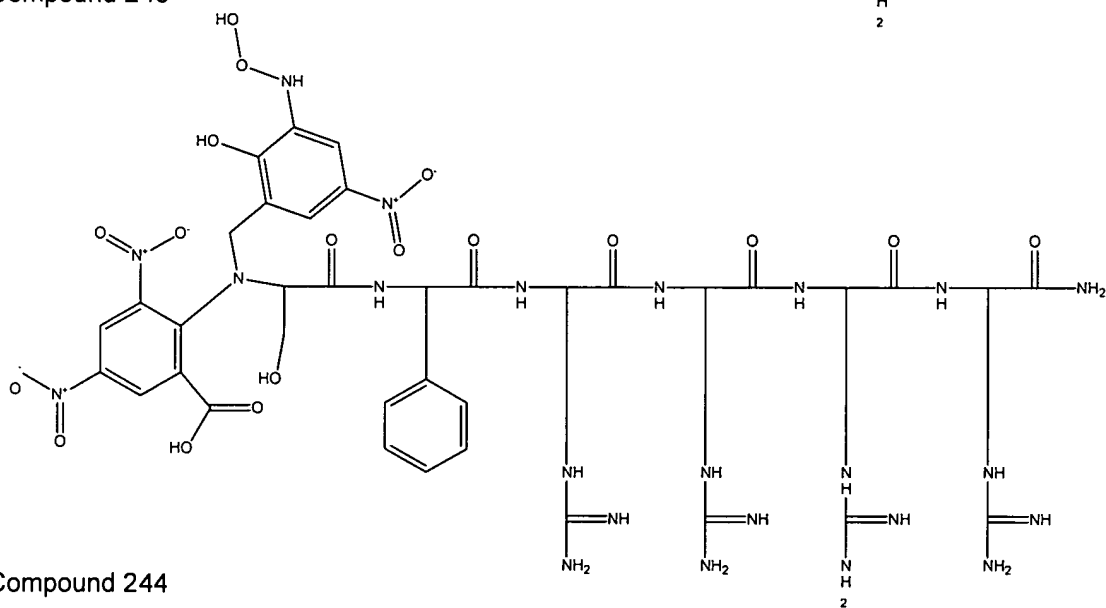
Compound 241



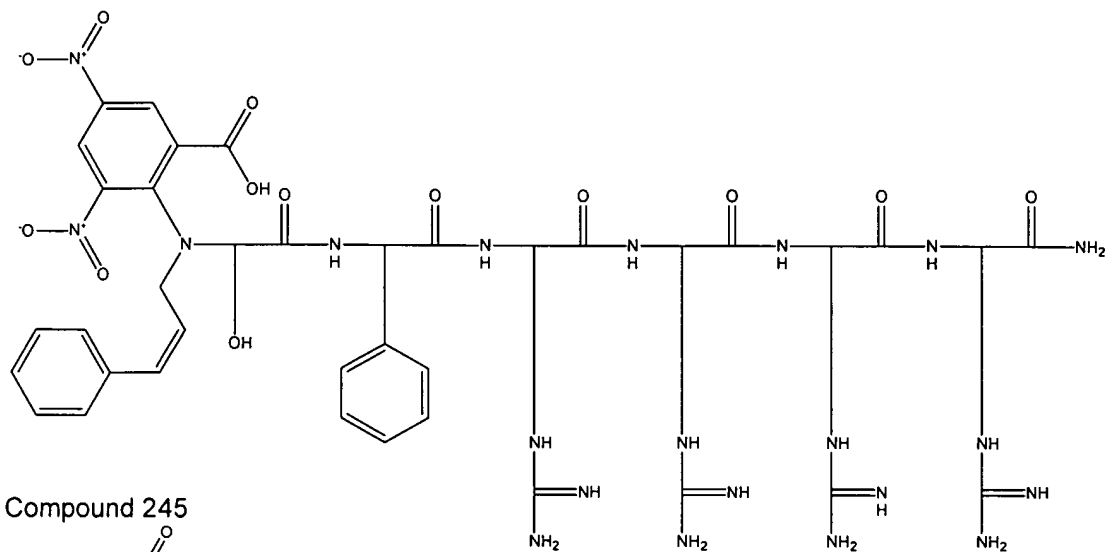
Compound 242



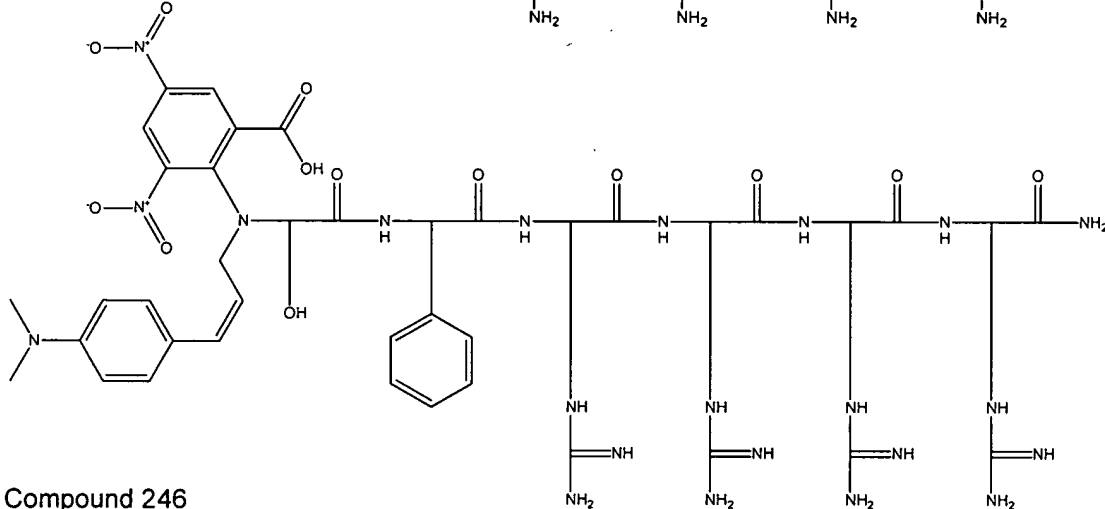
Compound 243



Compound 244

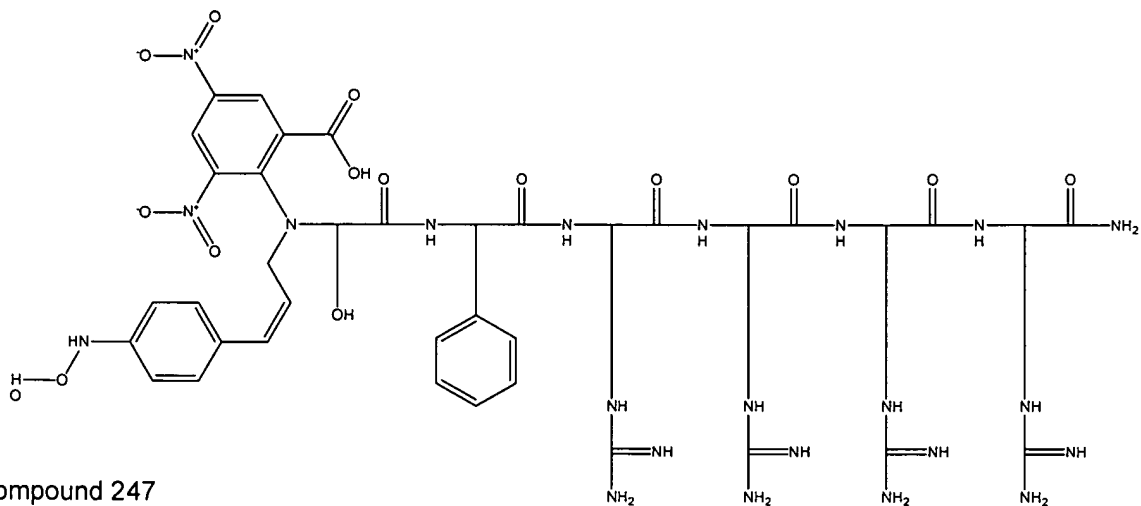


Compound 245

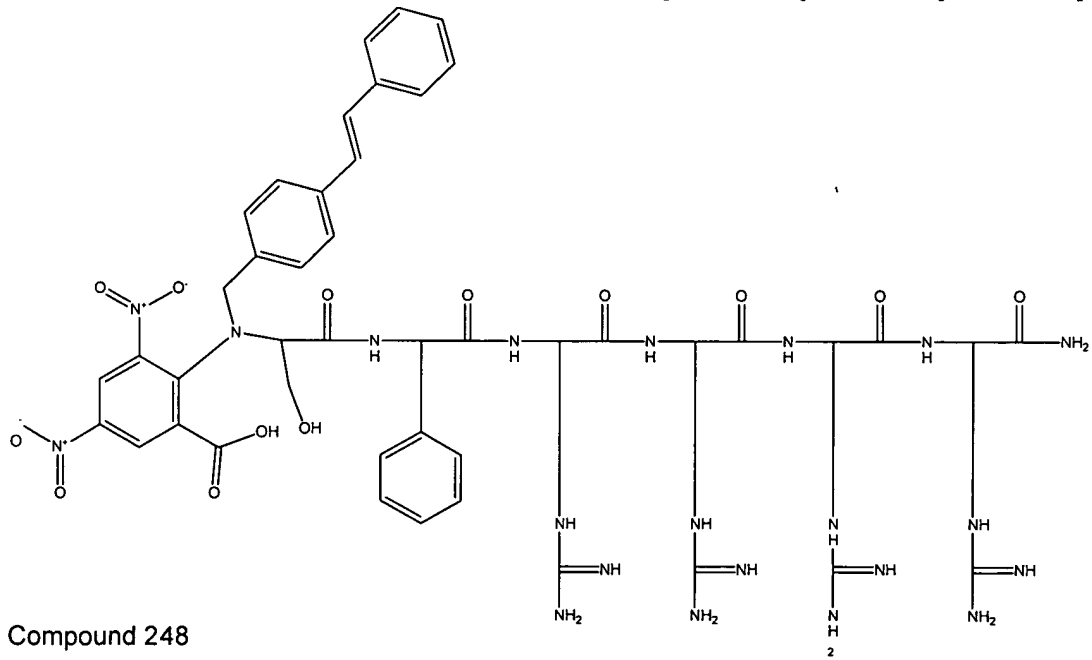


Compound 246

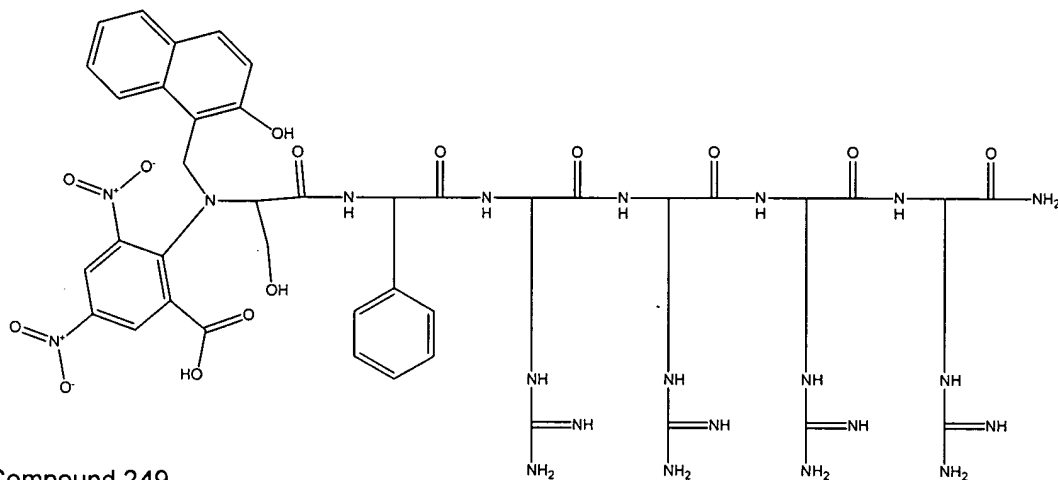
page 106 of 192



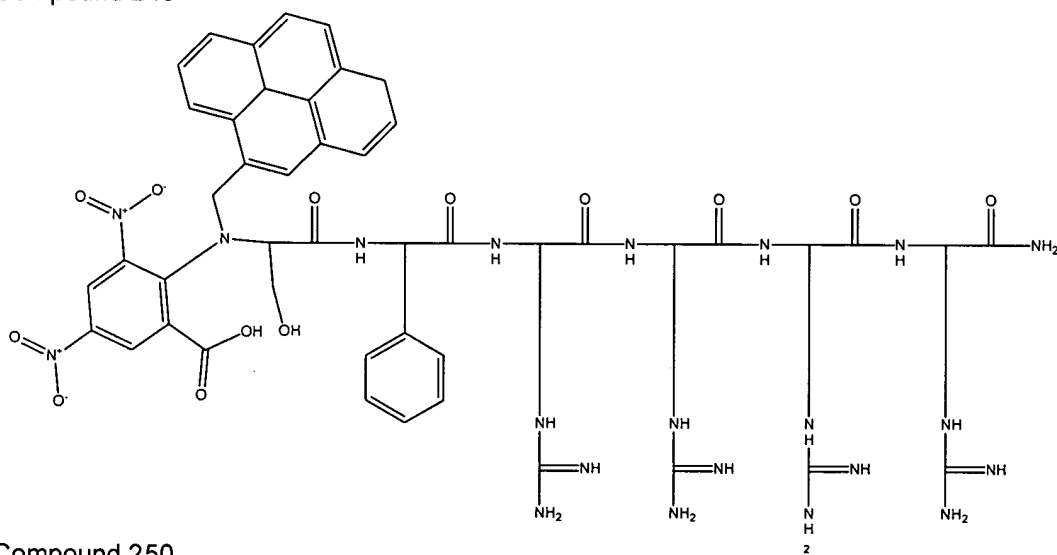
Compound 247



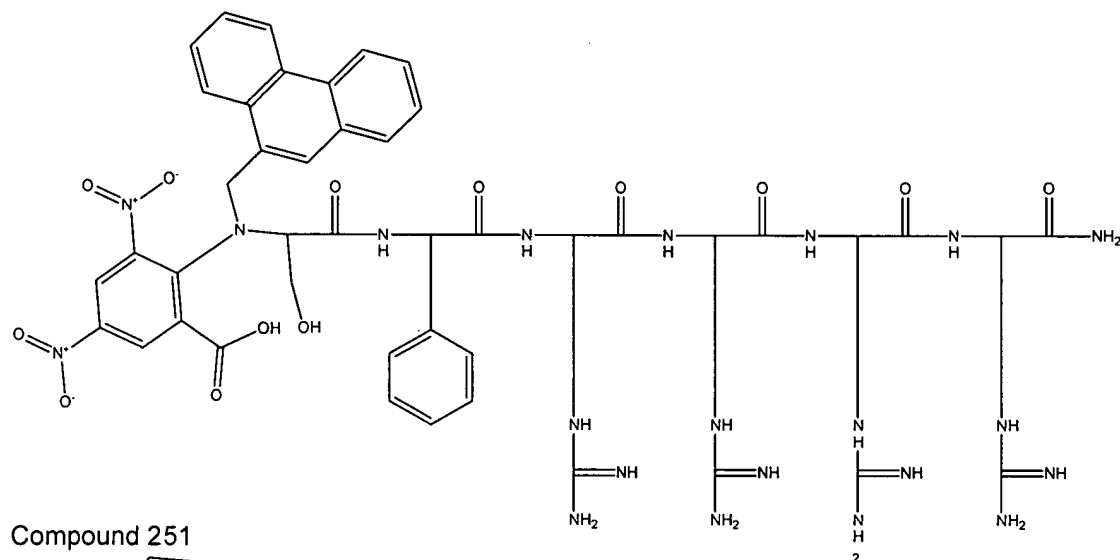
Compound 248



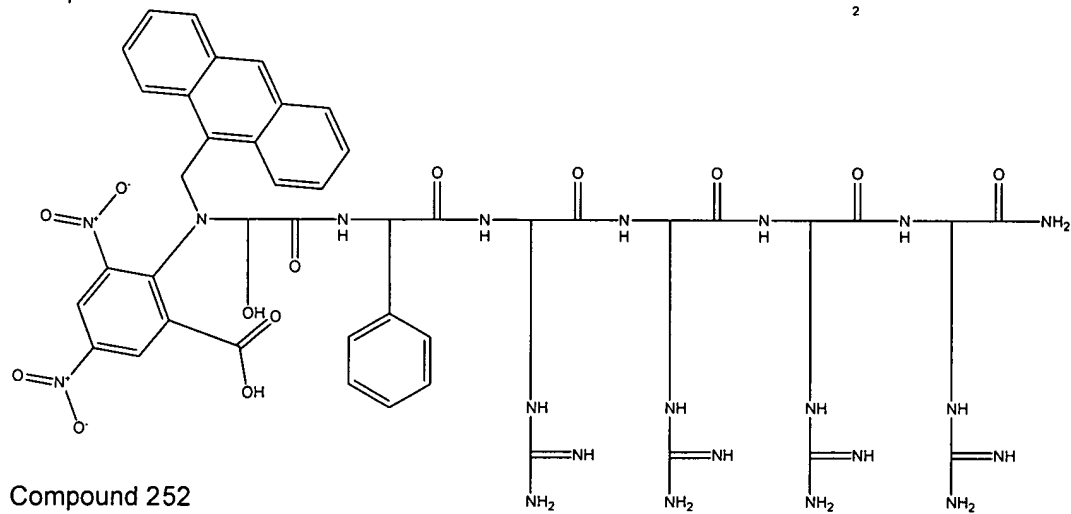
Compound 249



Compound 250

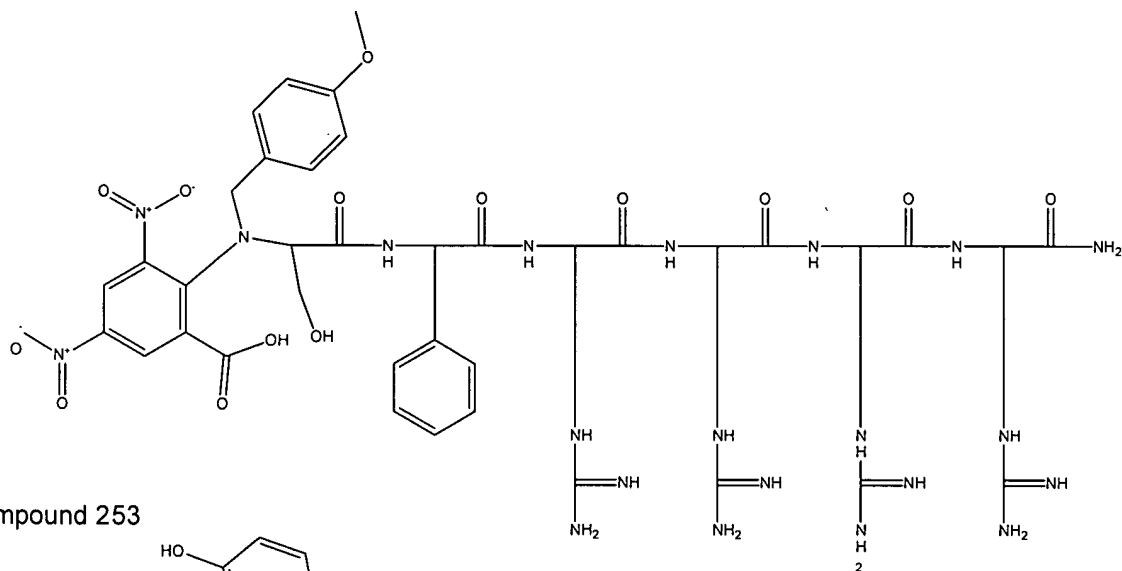


Compound 251

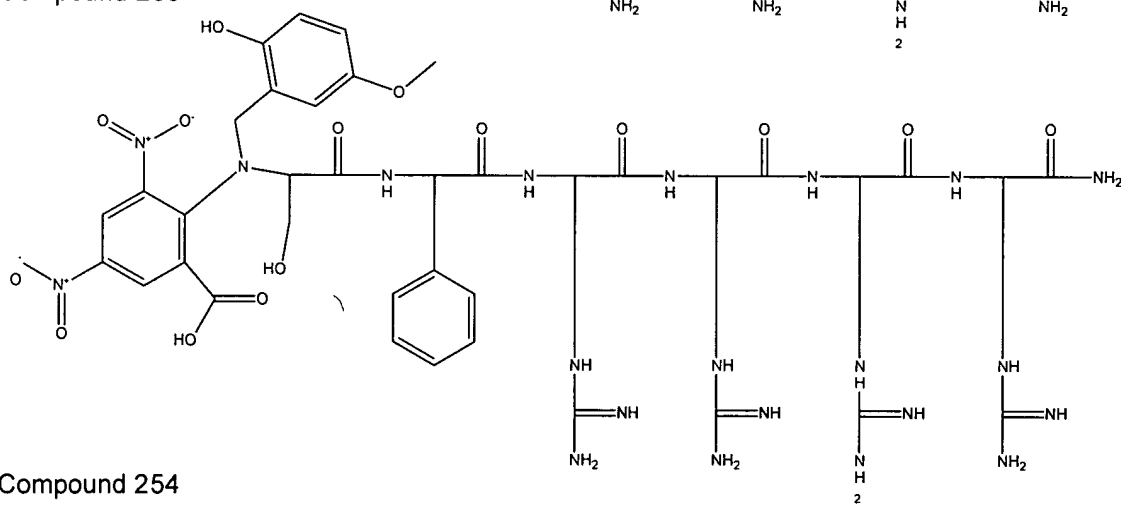


Compound 252

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 109 of 192

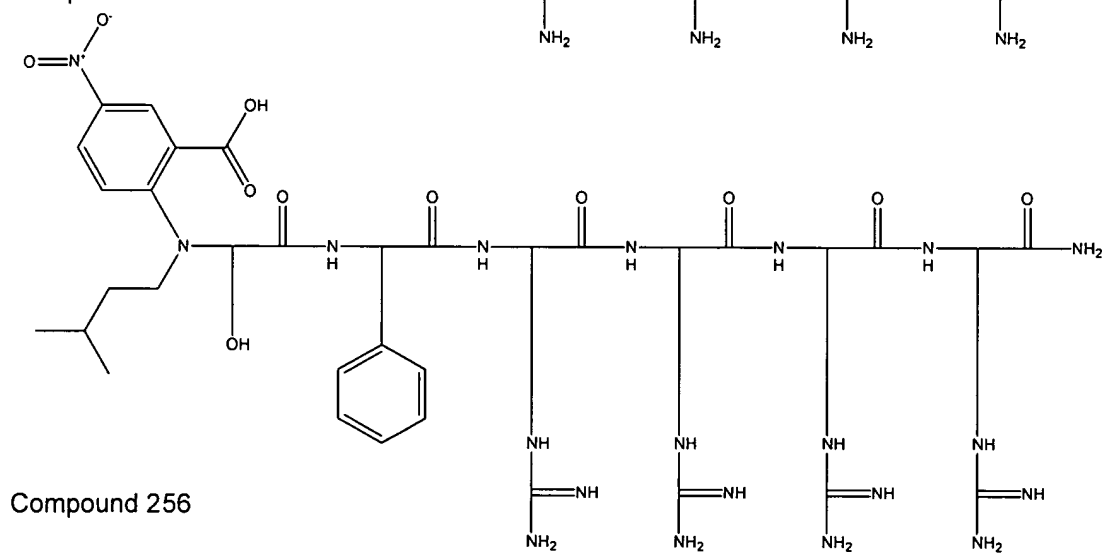
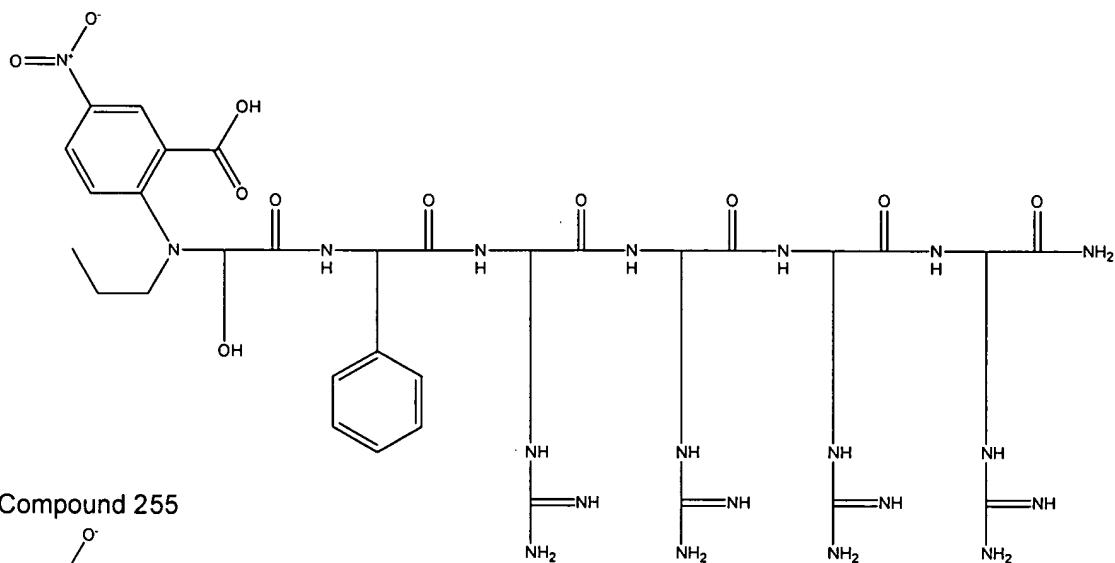


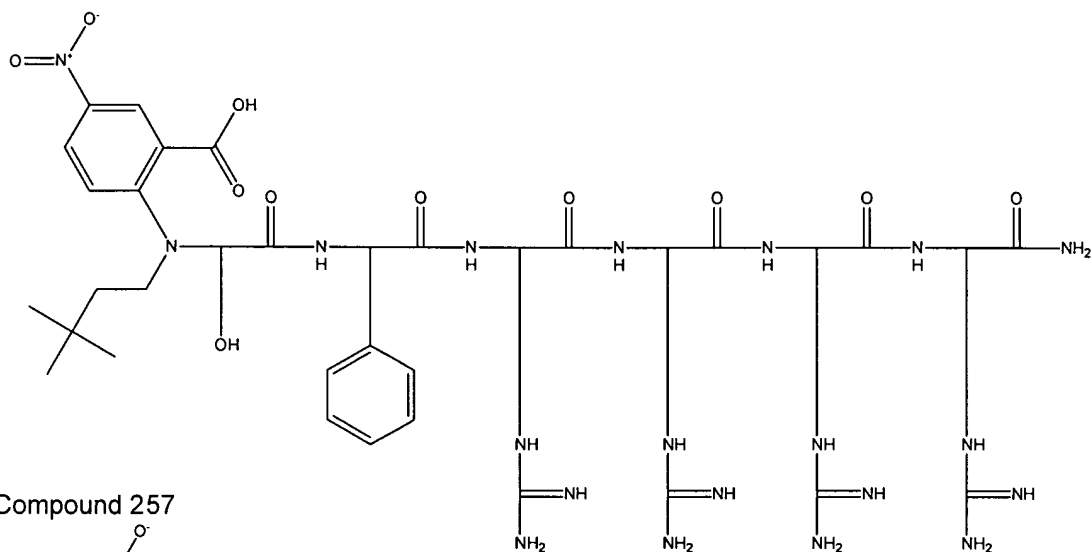
Compound 253



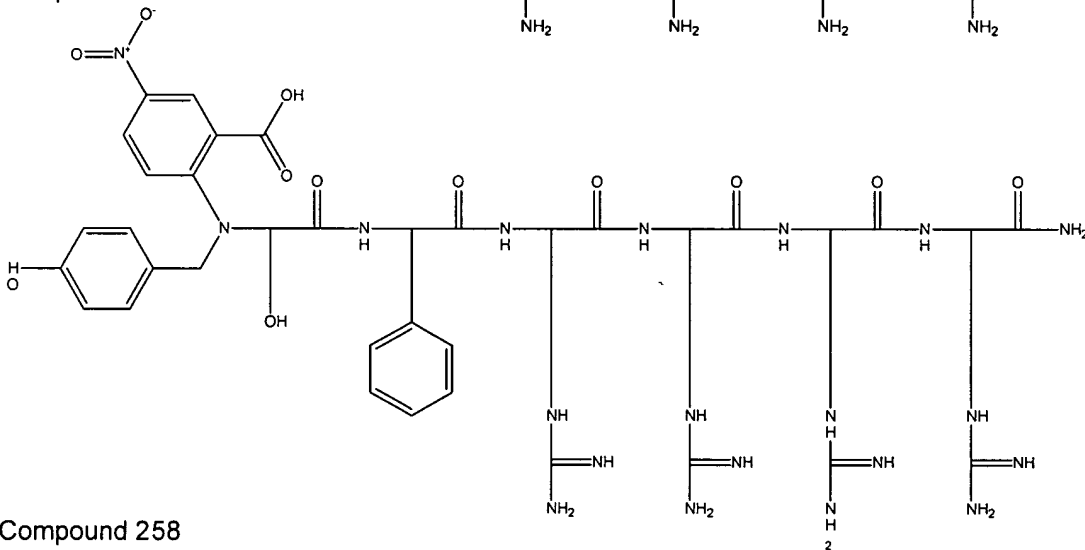
Compound 254

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 110 of 192





Compound 257



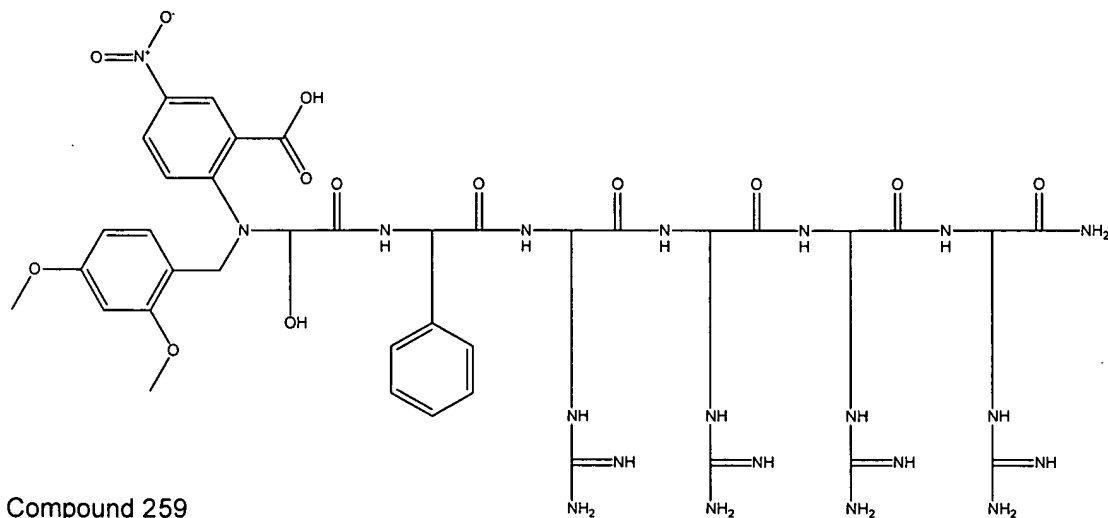
Compound 258

Applicant: David S. Lawrence

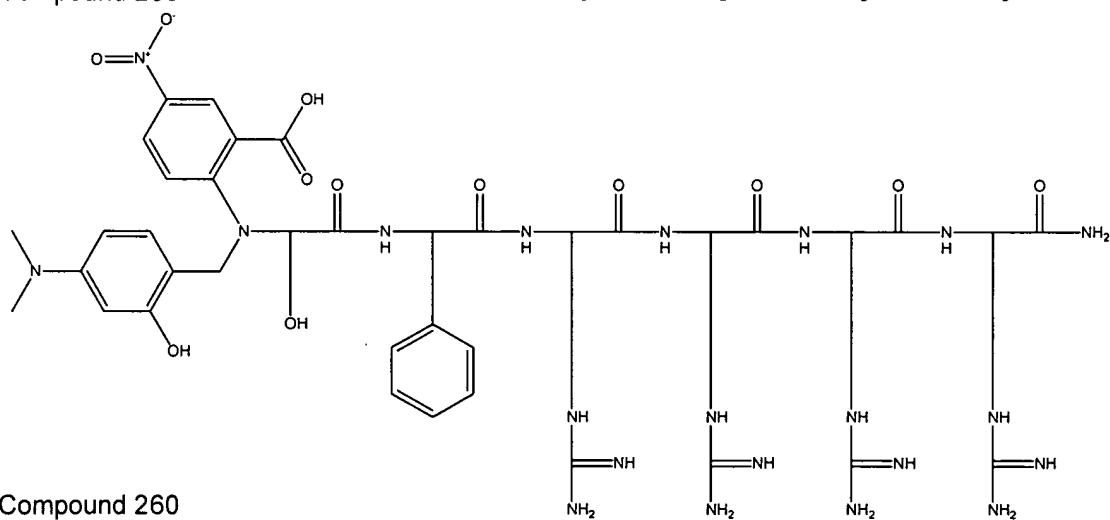
Serial No.: 10/755,086

Filed: January 9, 2004

page 112 of 192

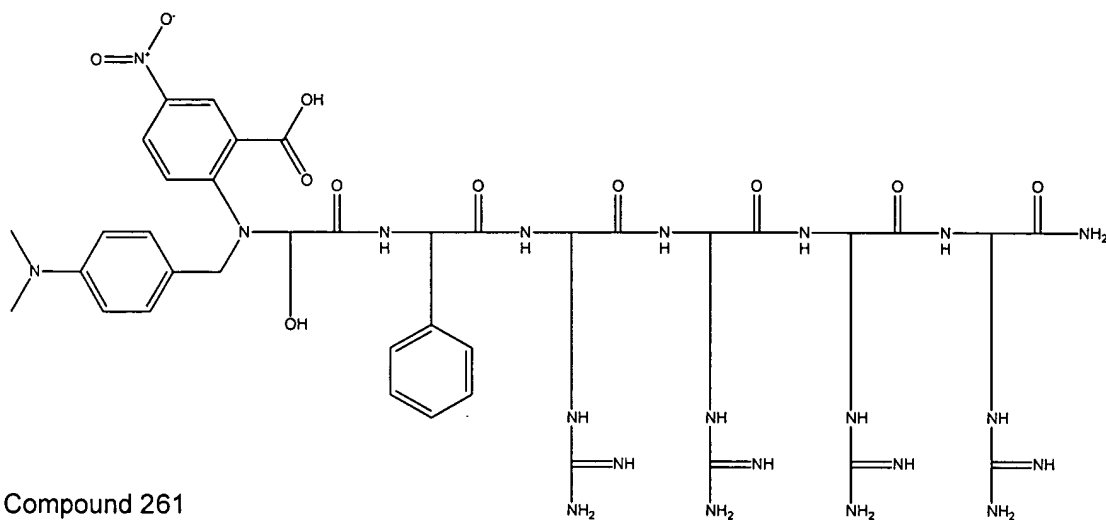


Compound 259

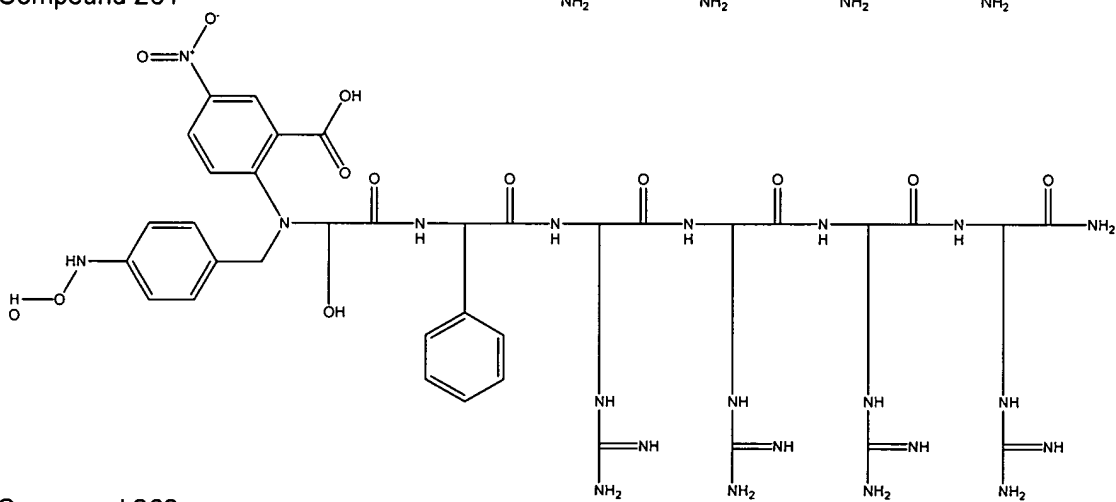


Compound 260

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 113 of 192

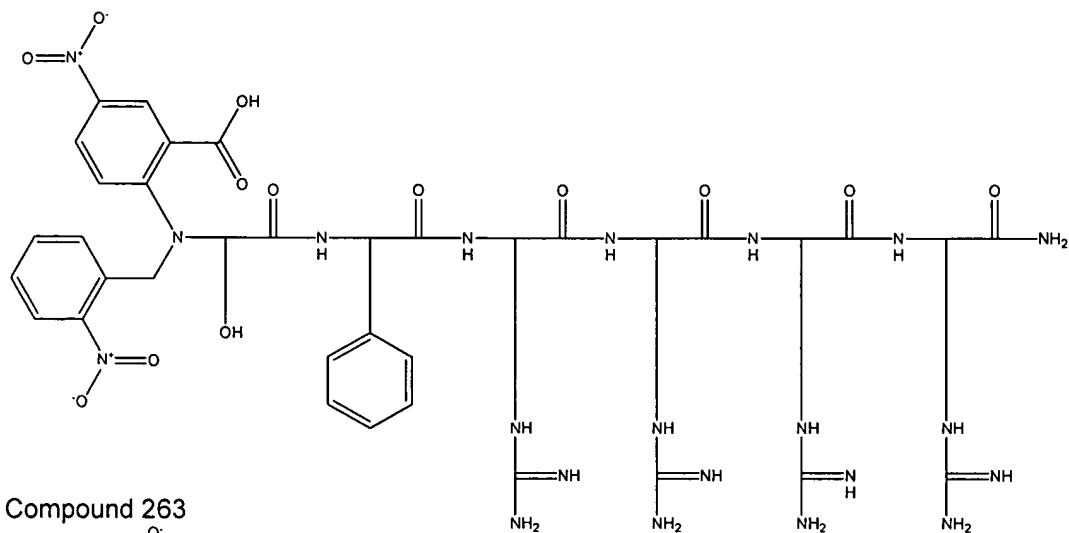


Compound 261

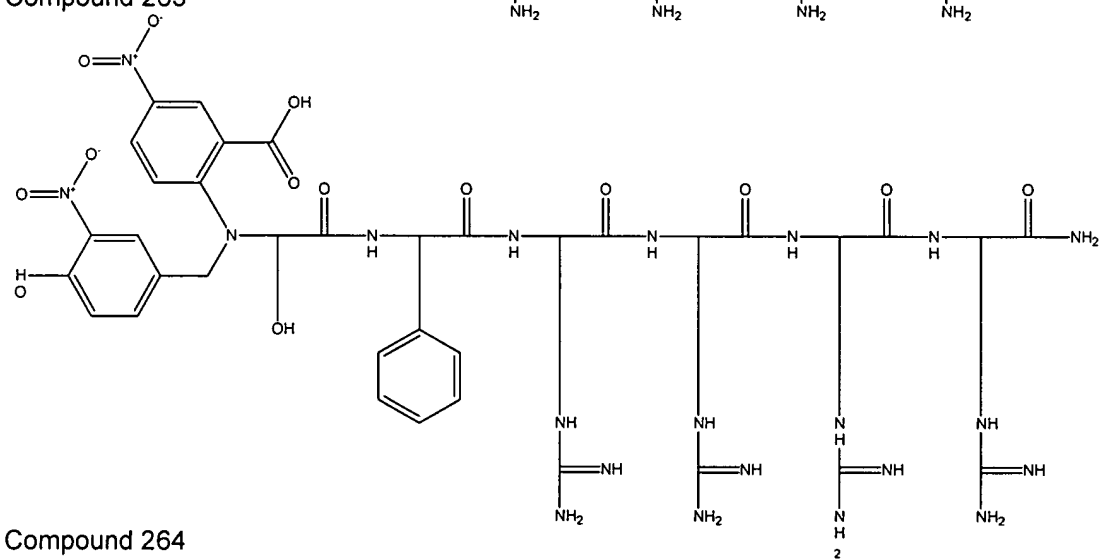


Compound 262

Applicant: David S. Lawrence
 Serial No.: 10/755,086
 Filed: January 9, 2004
 page 114 of 192

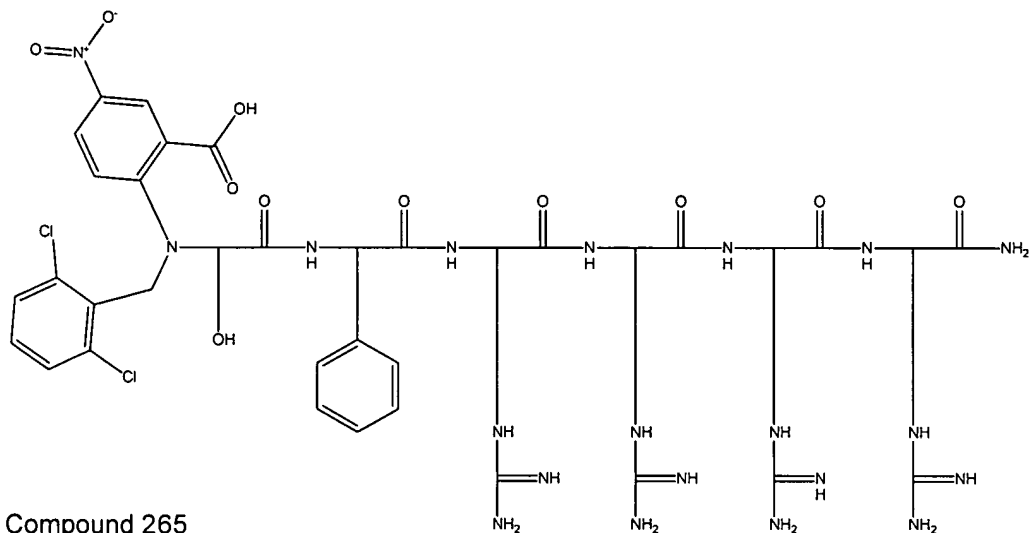


Compound 263

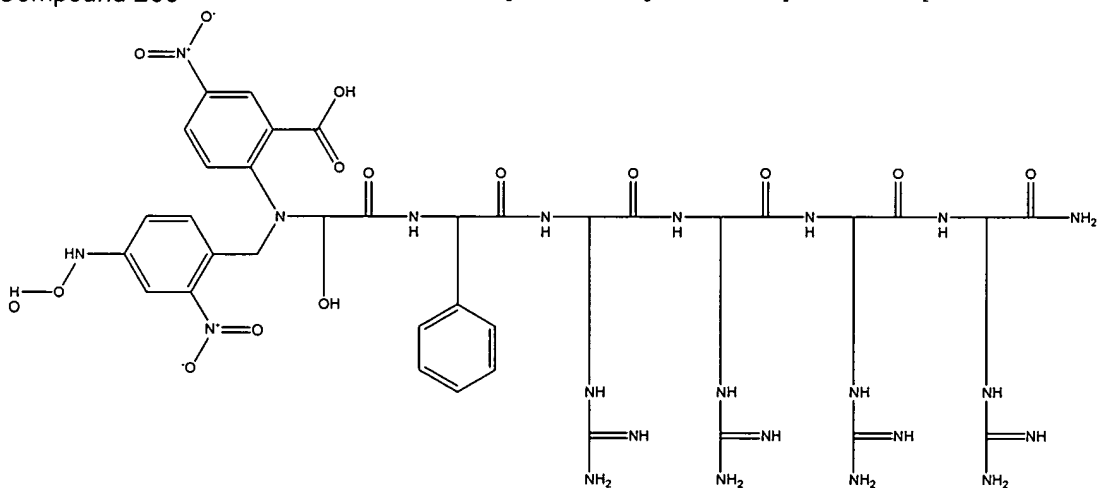


Compound 264

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 115 of 192

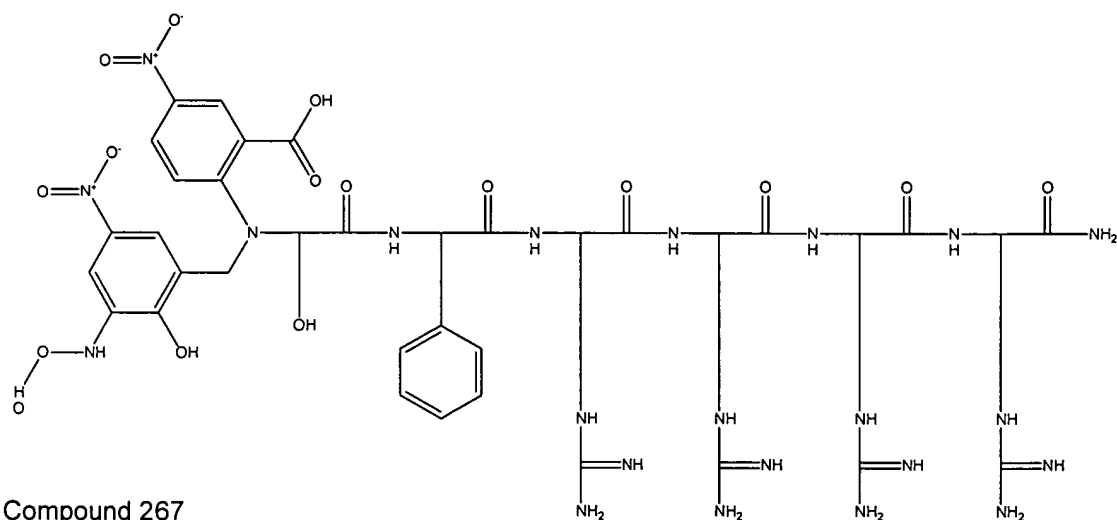


Compound 265

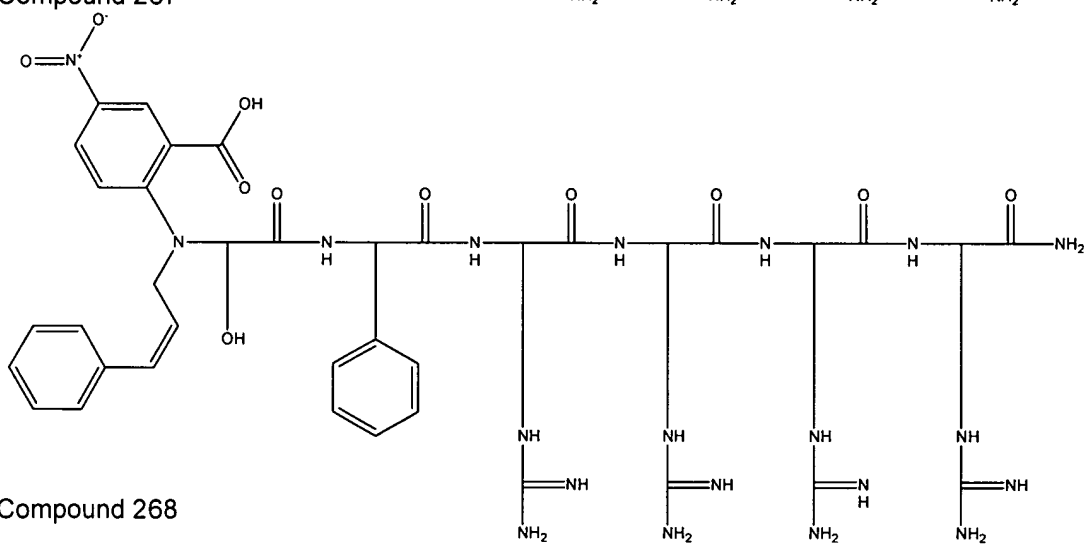


Compound 266

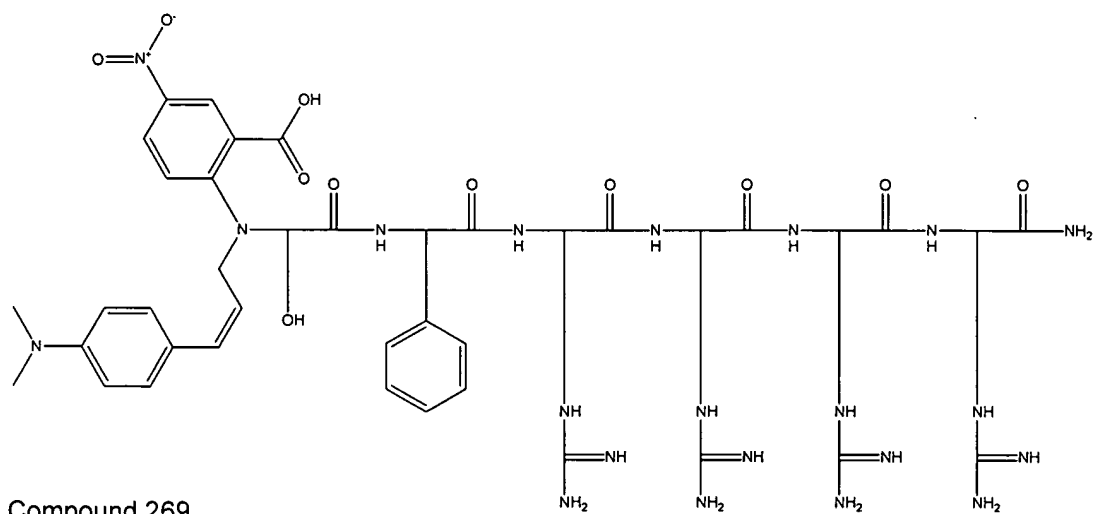
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 116 of 192



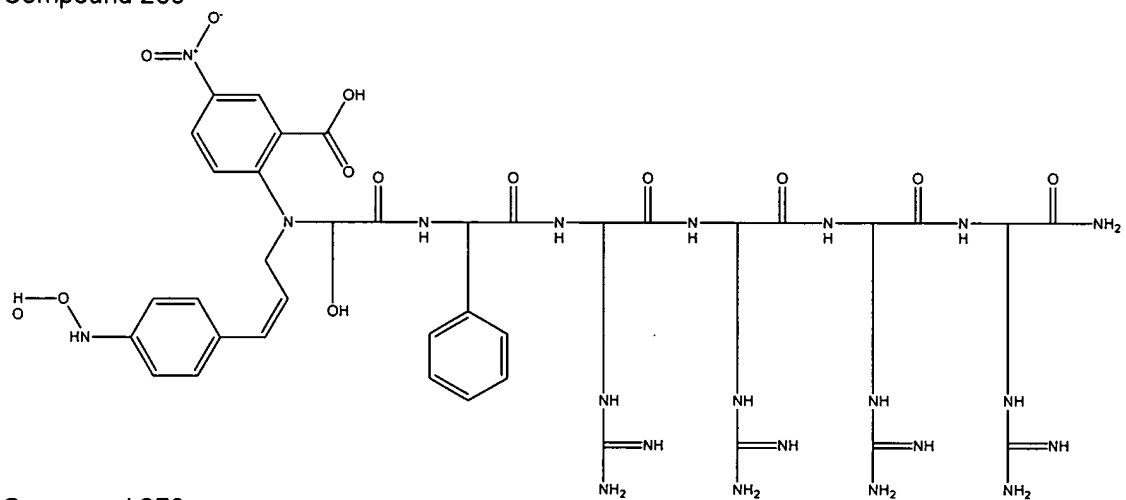
Compound 267



Compound 268

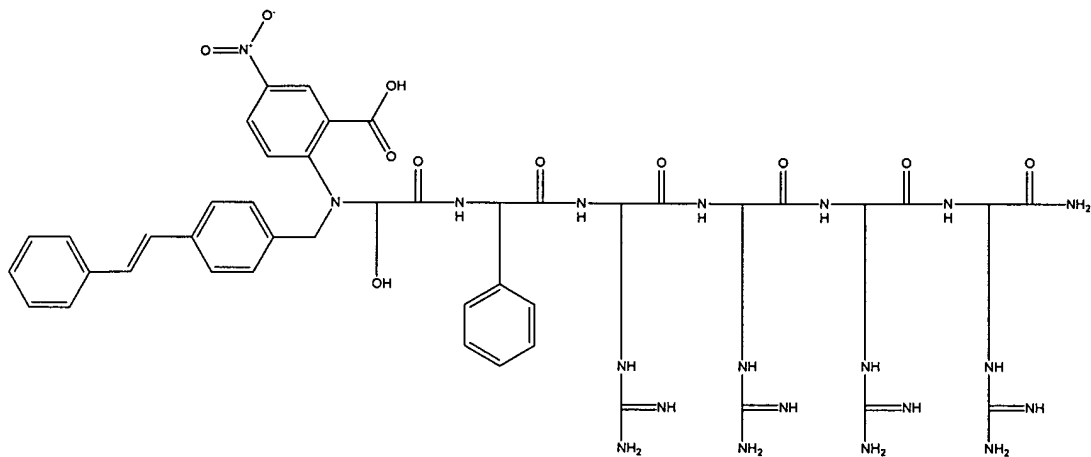


Compound 269

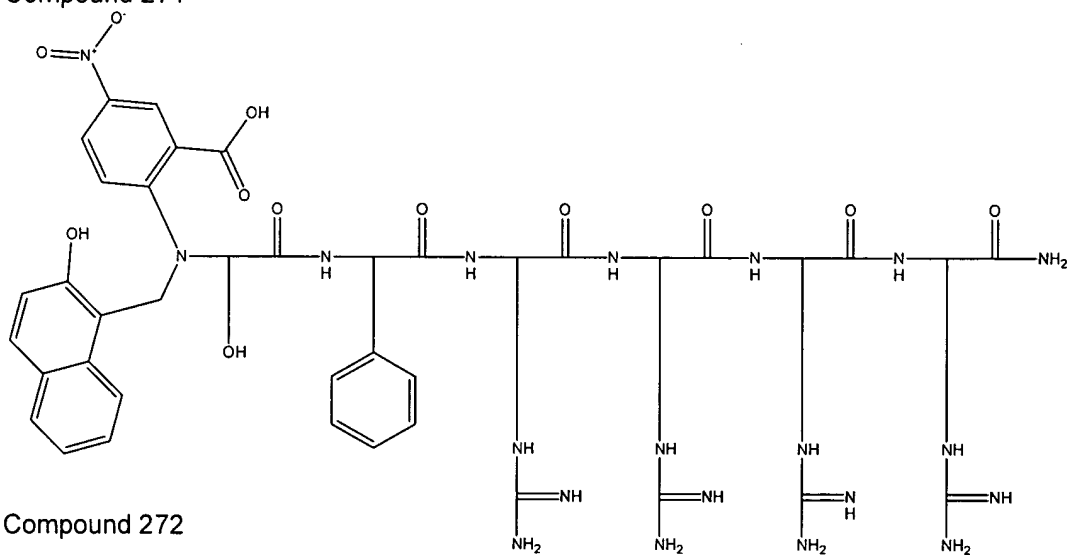


Compound 270

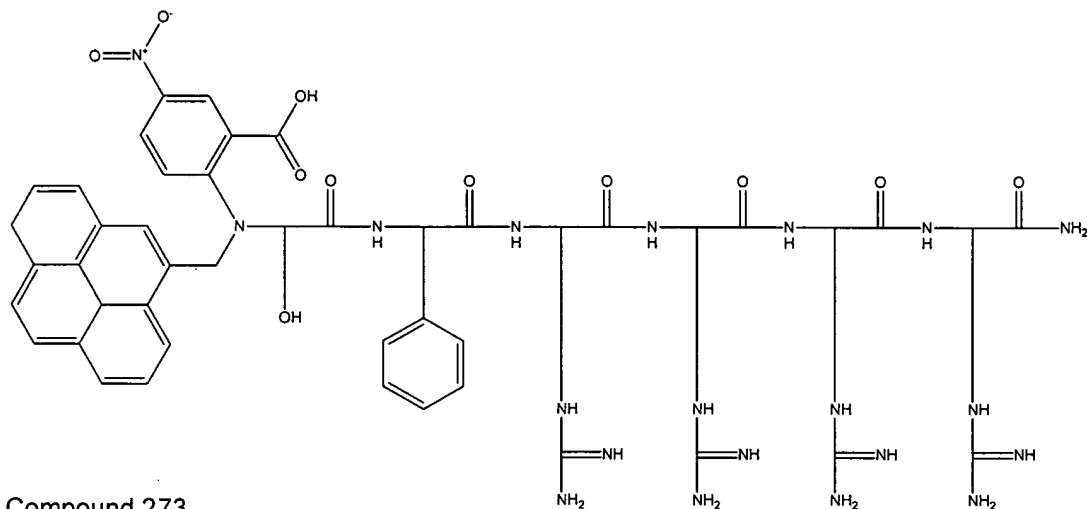
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 118 of 192



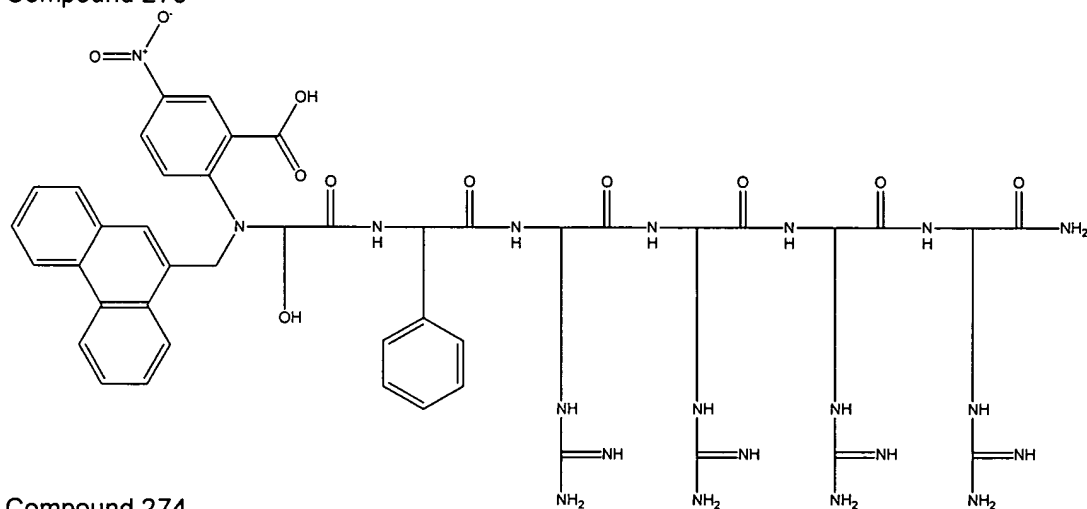
Compound 271



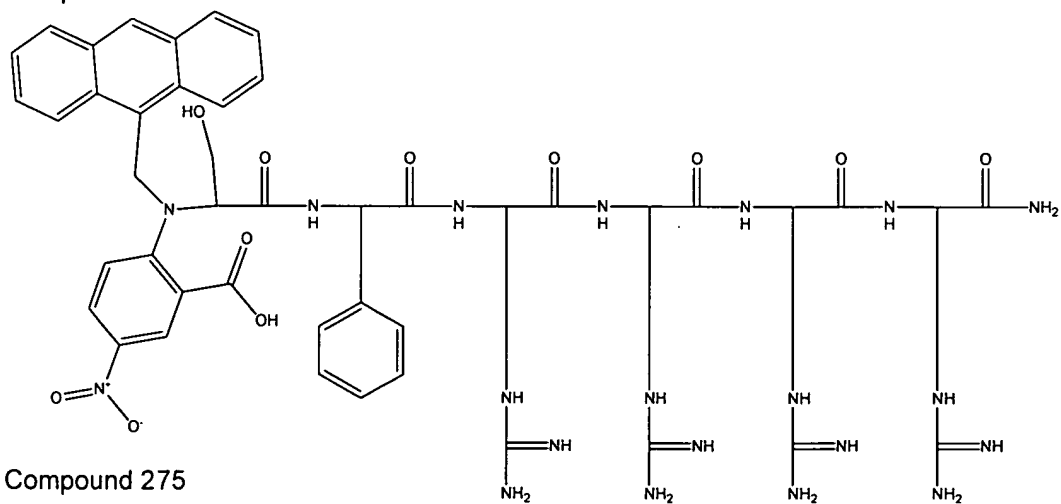
Compound 272



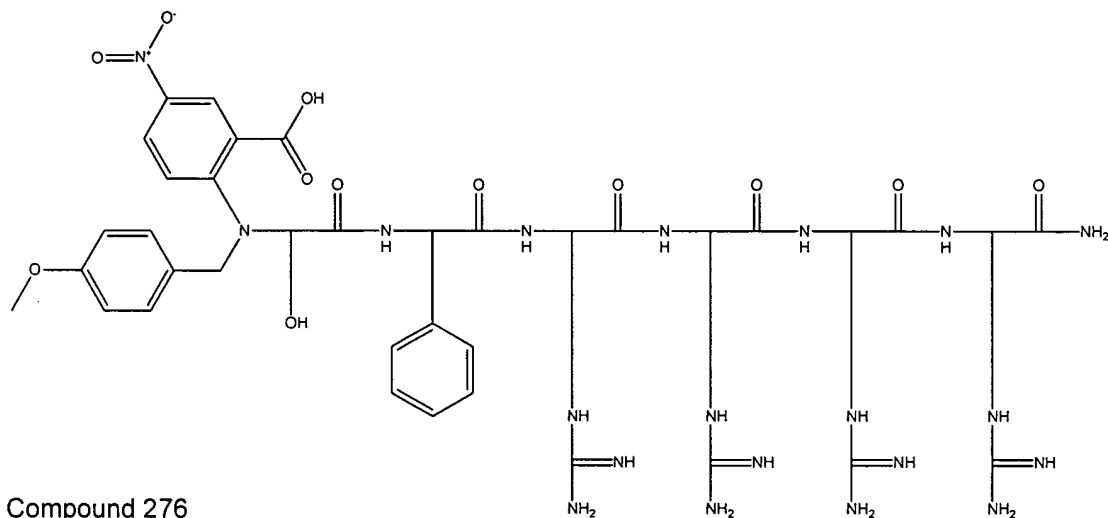
Compound 273



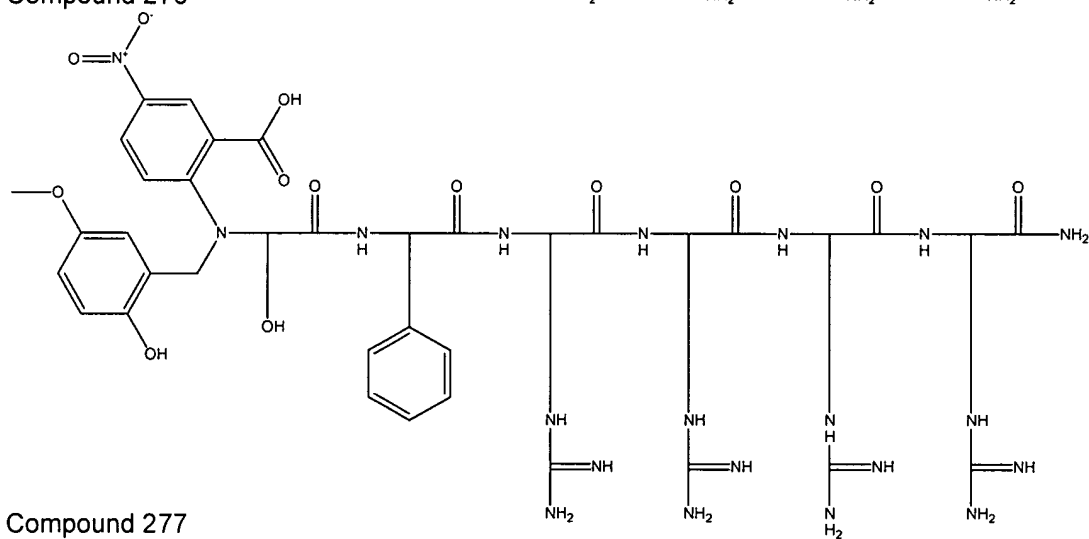
Compound 274



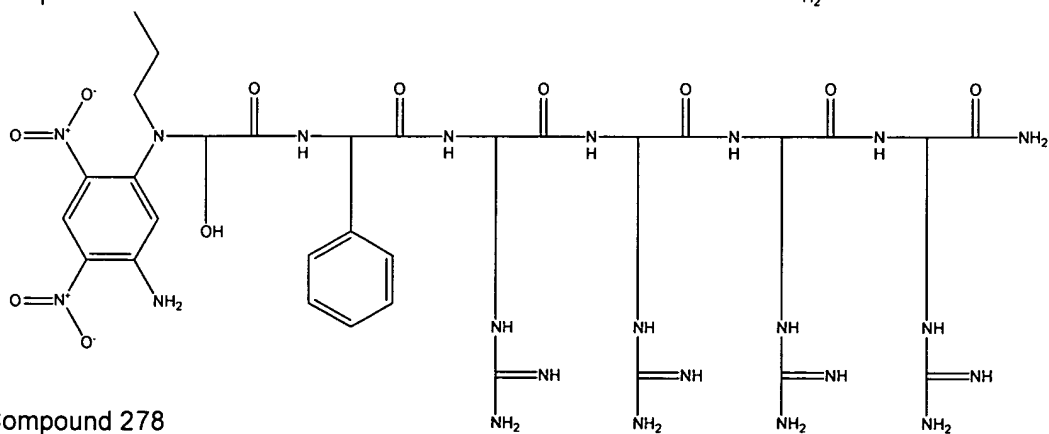
Compound 275



Compound 276



Compound 277



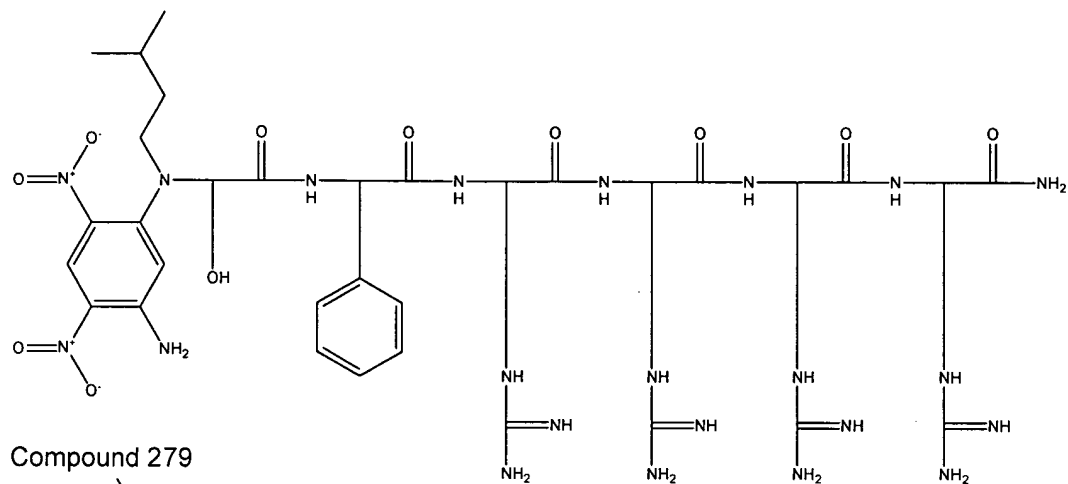
Compound 278

Applicant: David S. Lawrence

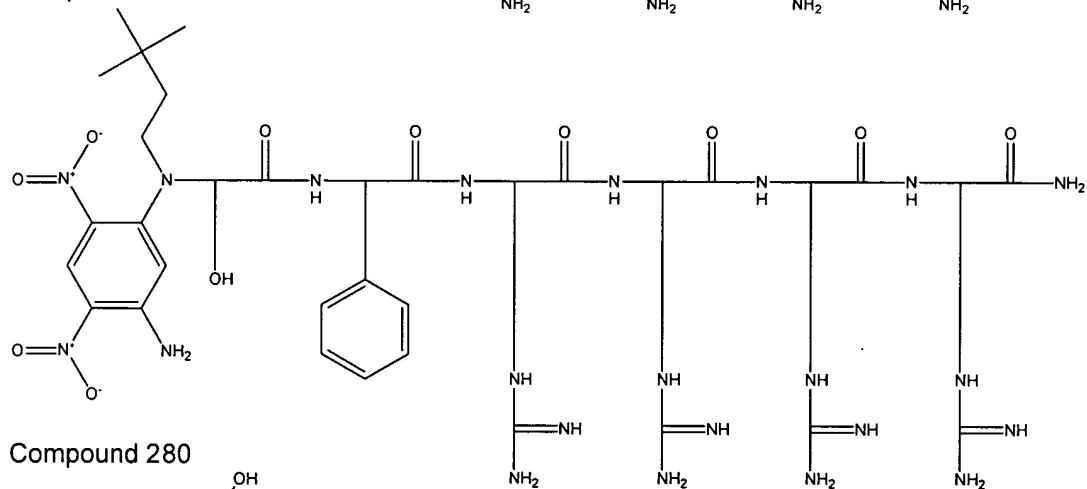
Serial No.: 10/755,086

Filed: January 9, 2004

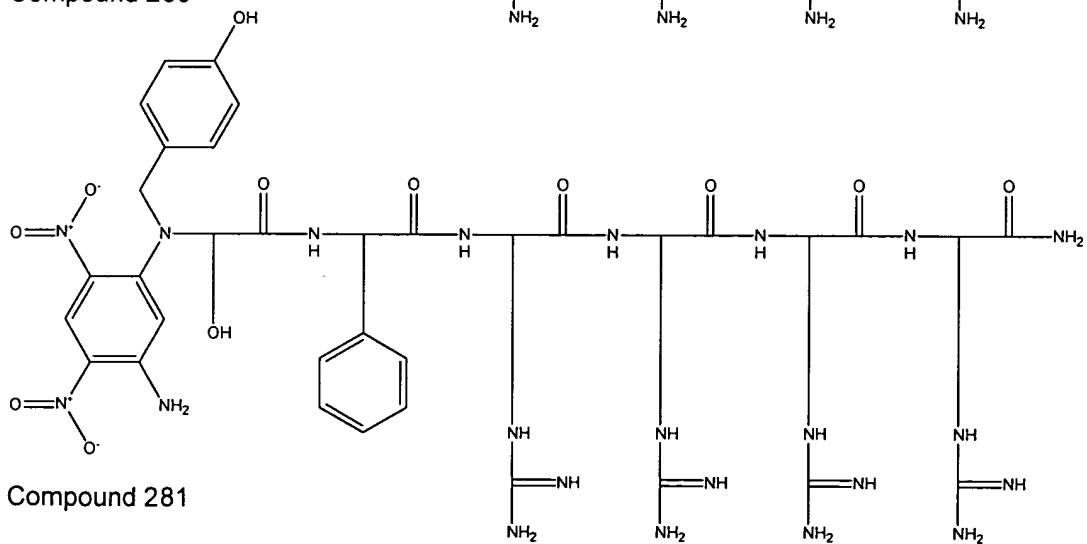
page 121 of 192



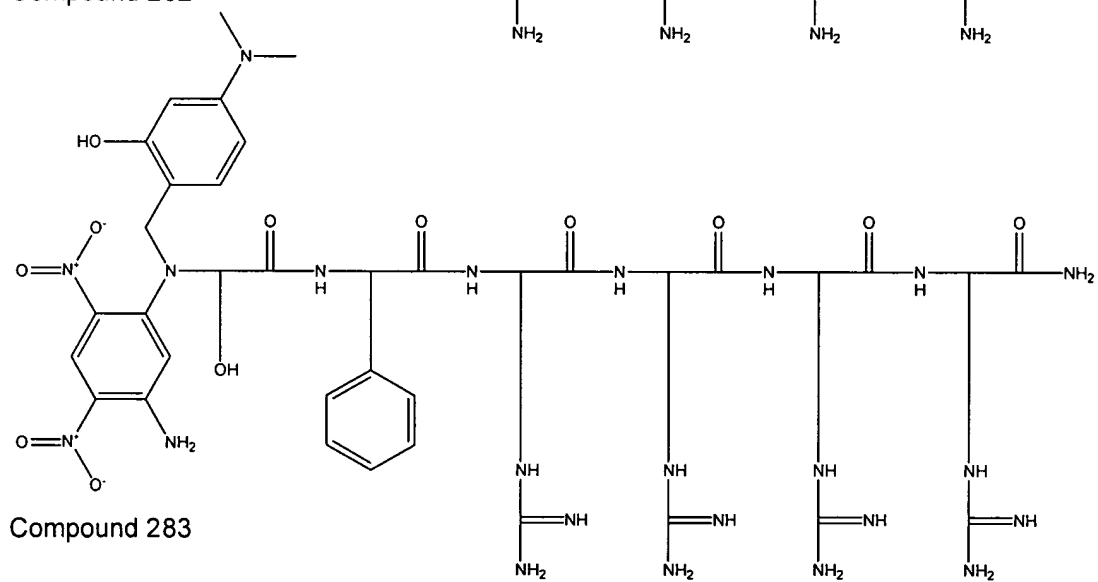
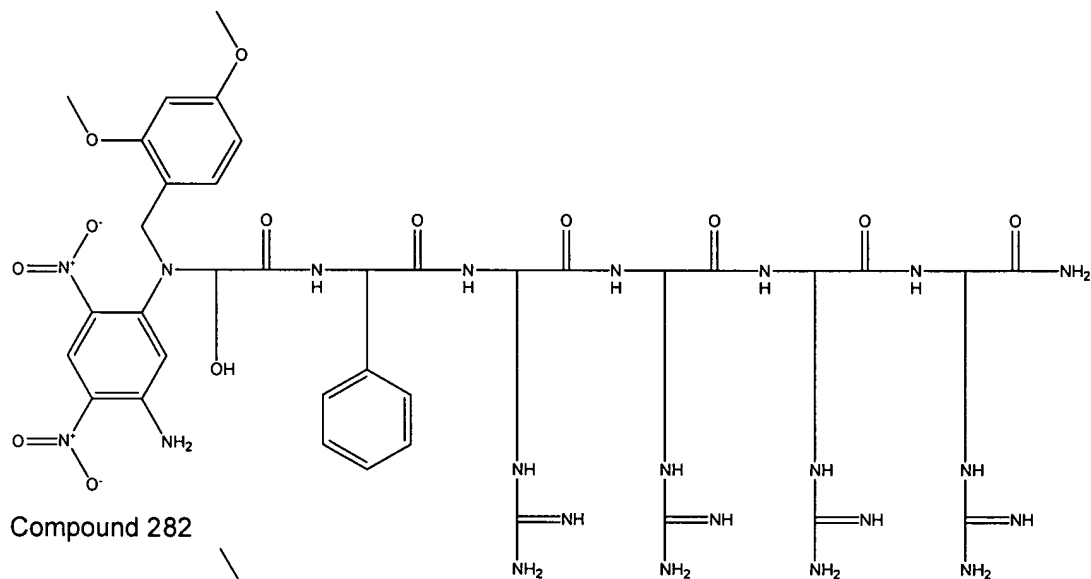
Compound 279

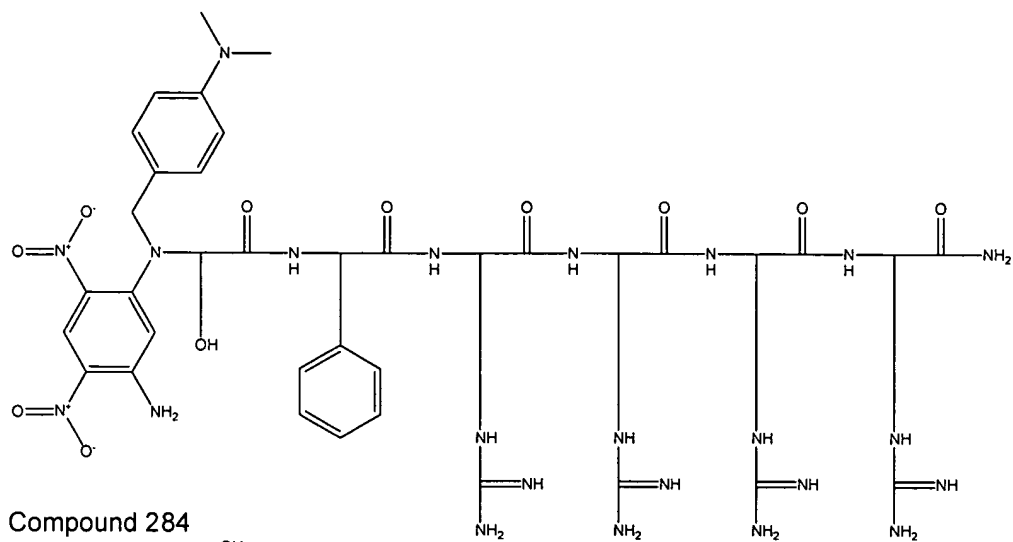


Compound 280

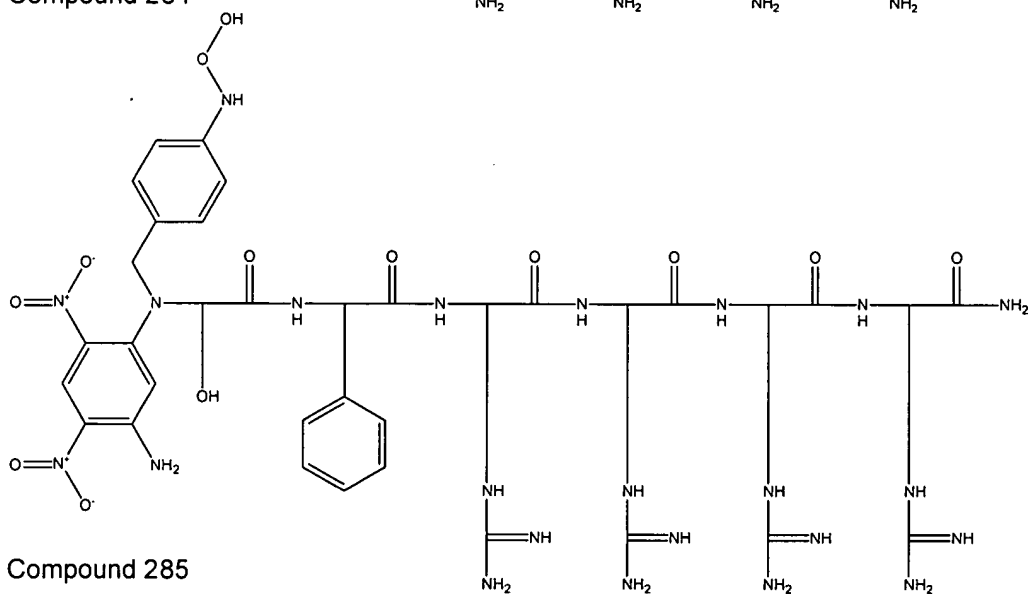


Compound 281

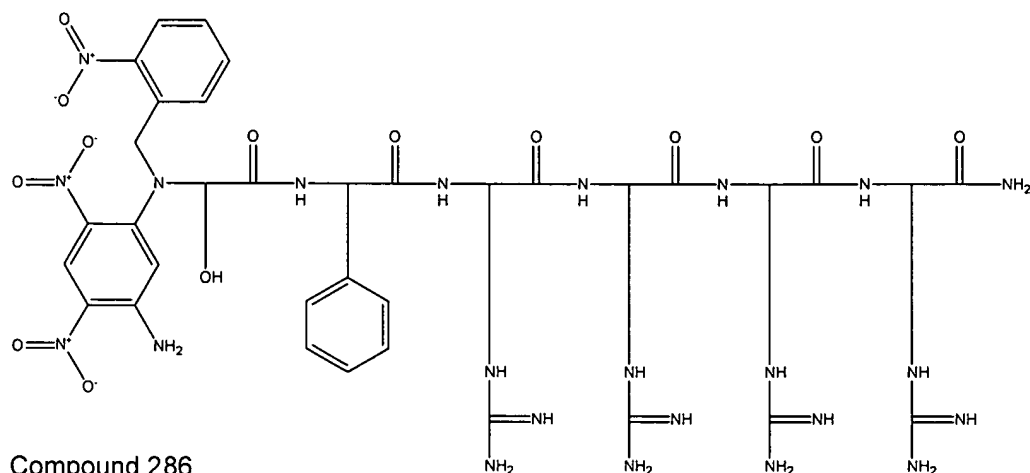




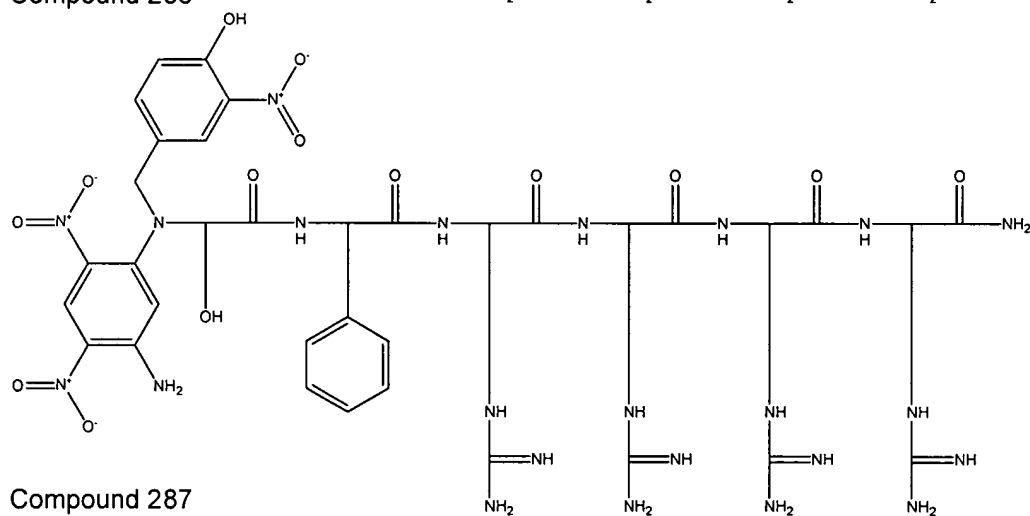
Compound 284



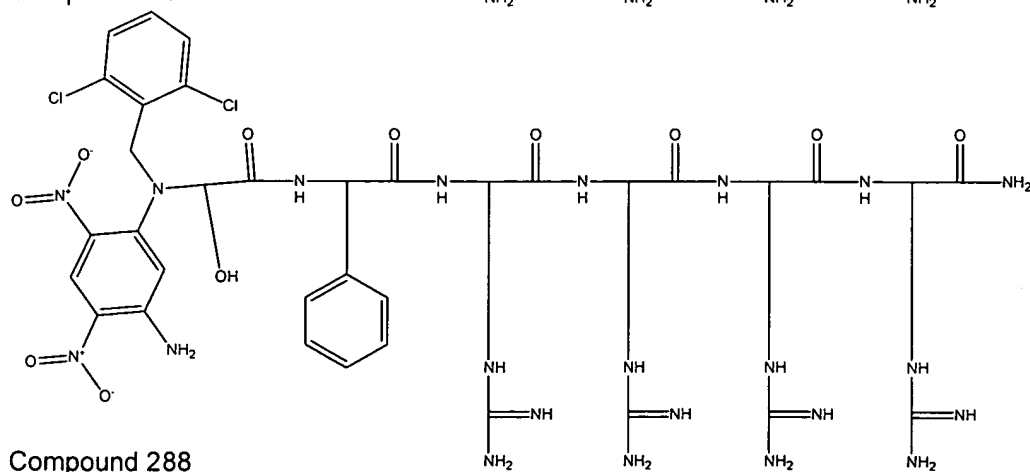
Compound 285



Compound 286

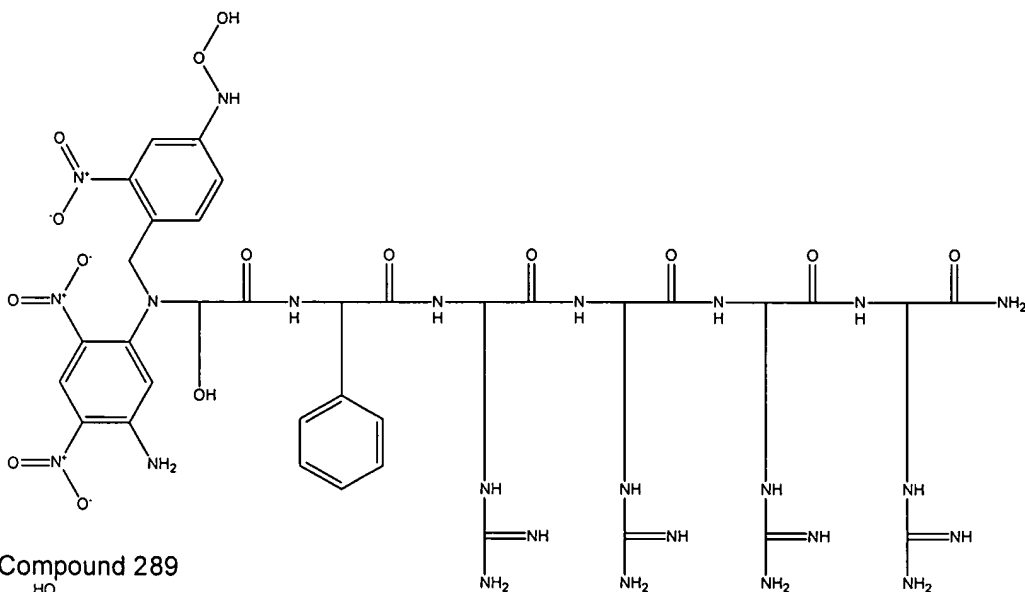


Compound 287

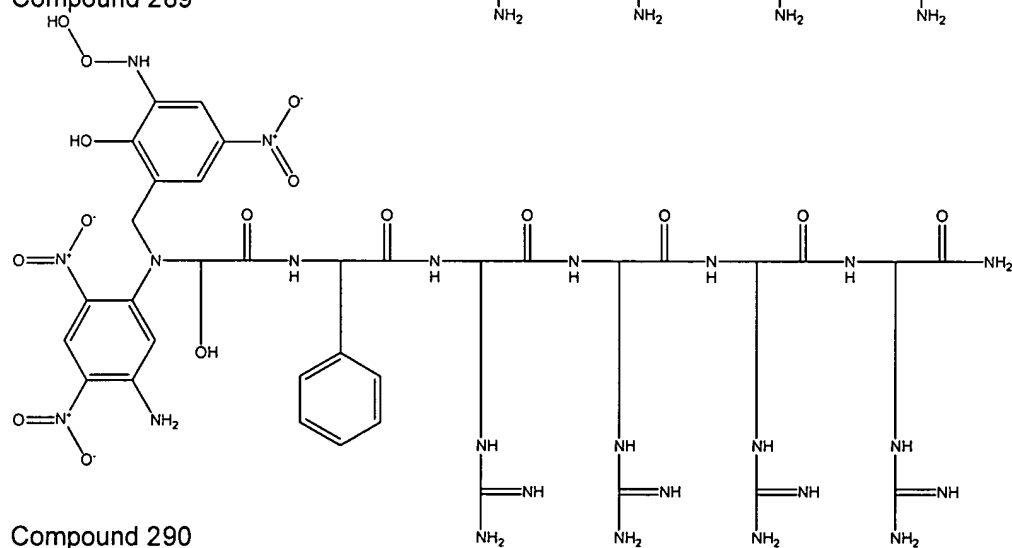


Compound 288

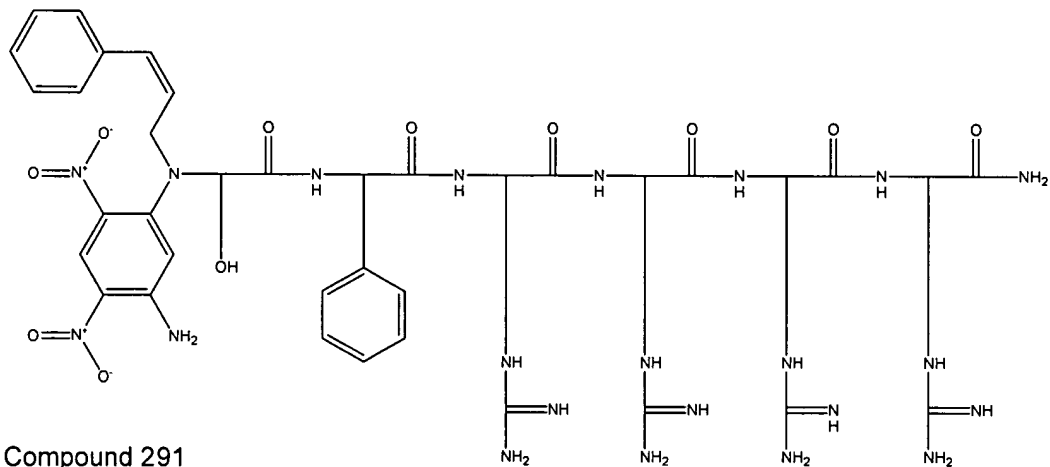
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 125 of 192



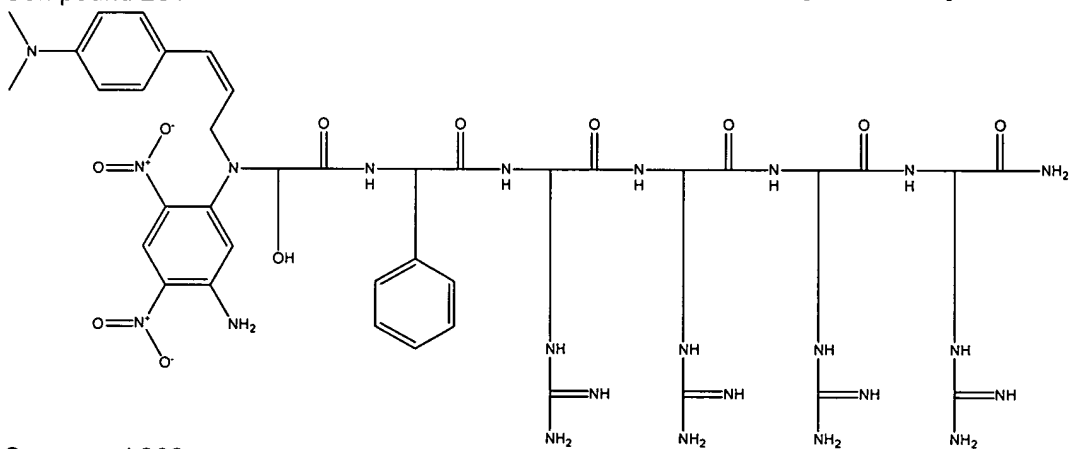
Compound 289



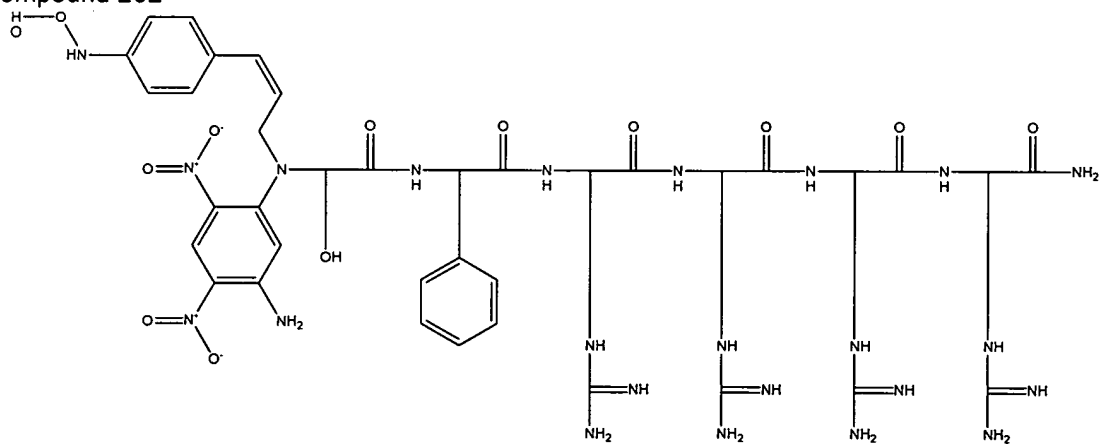
Compound 290



Compound 291

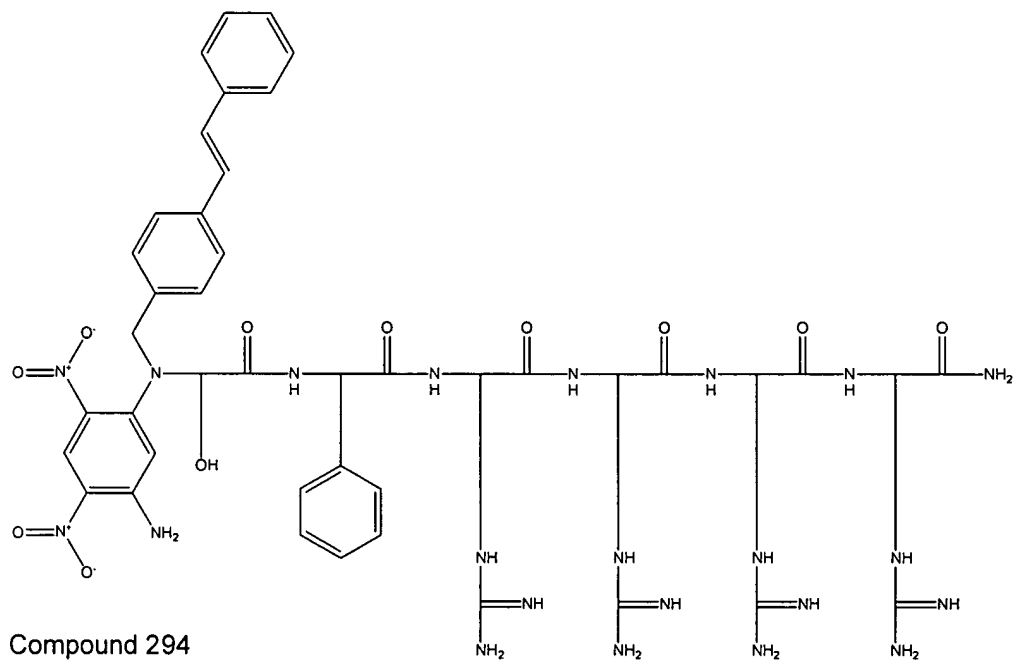


Compound 292

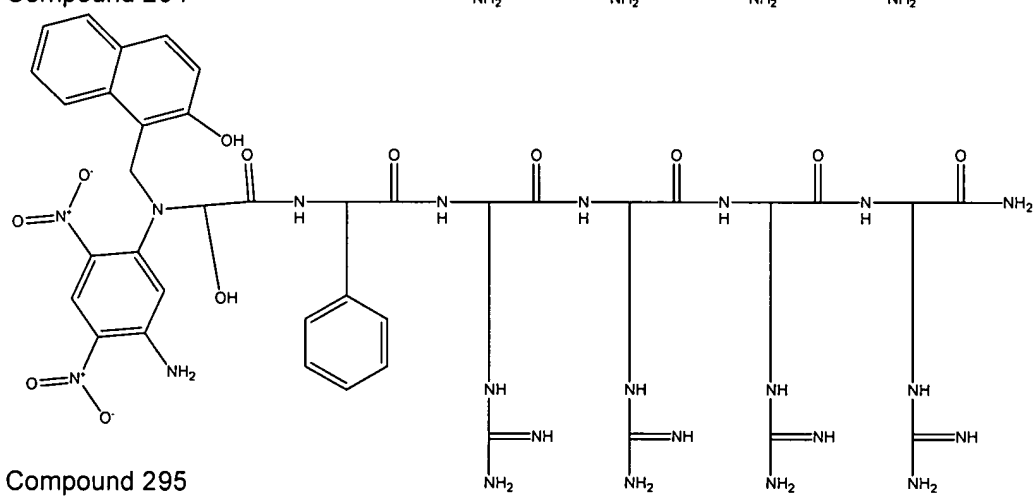


Compound 293

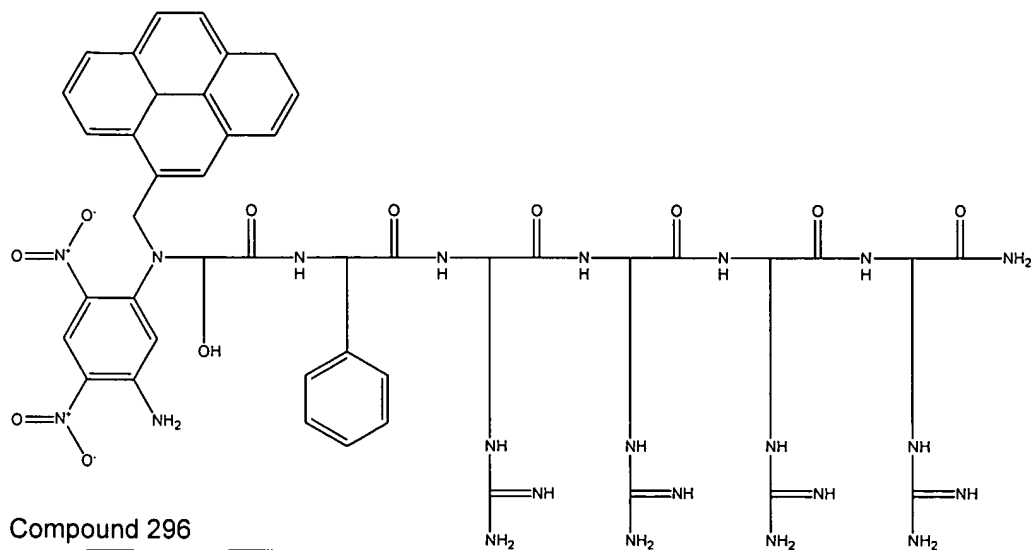
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 127 of 192



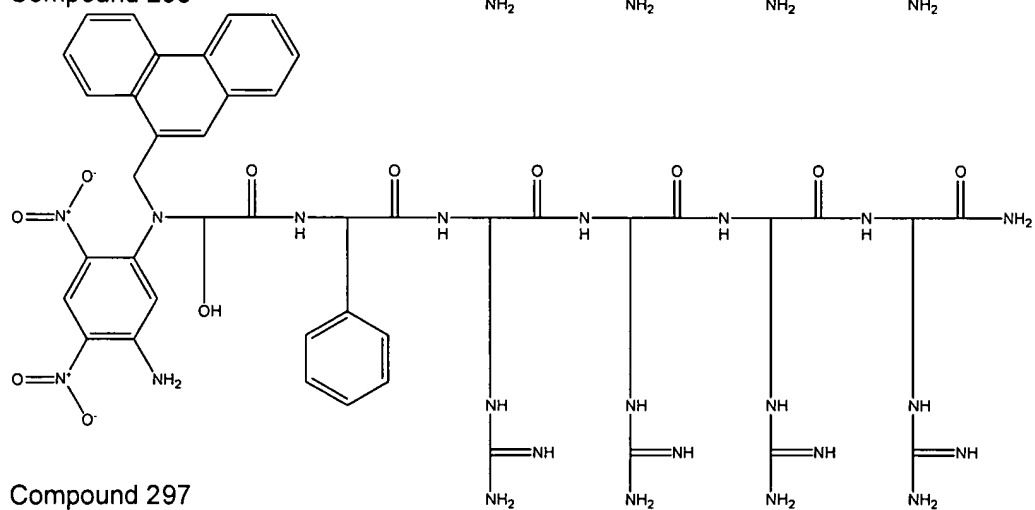
Compound 294



Compound 295

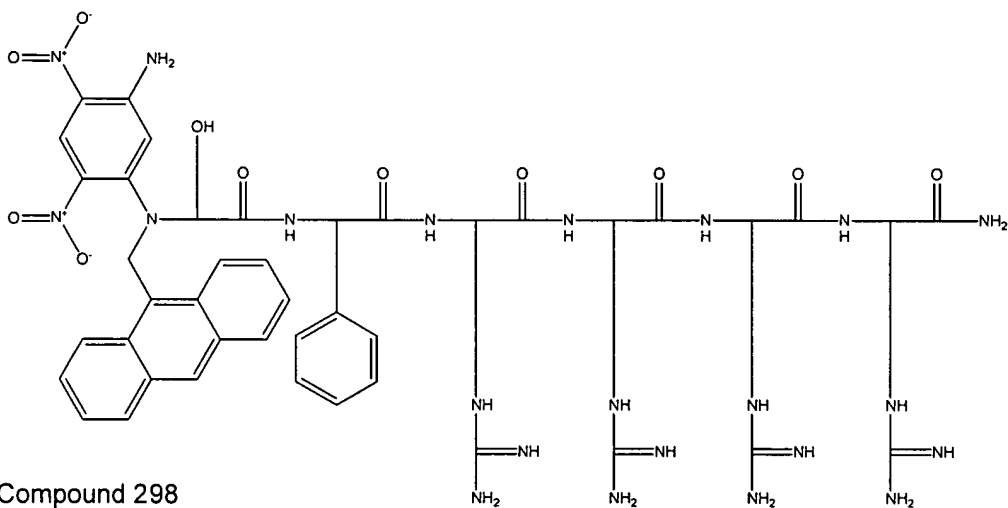


Compound 296

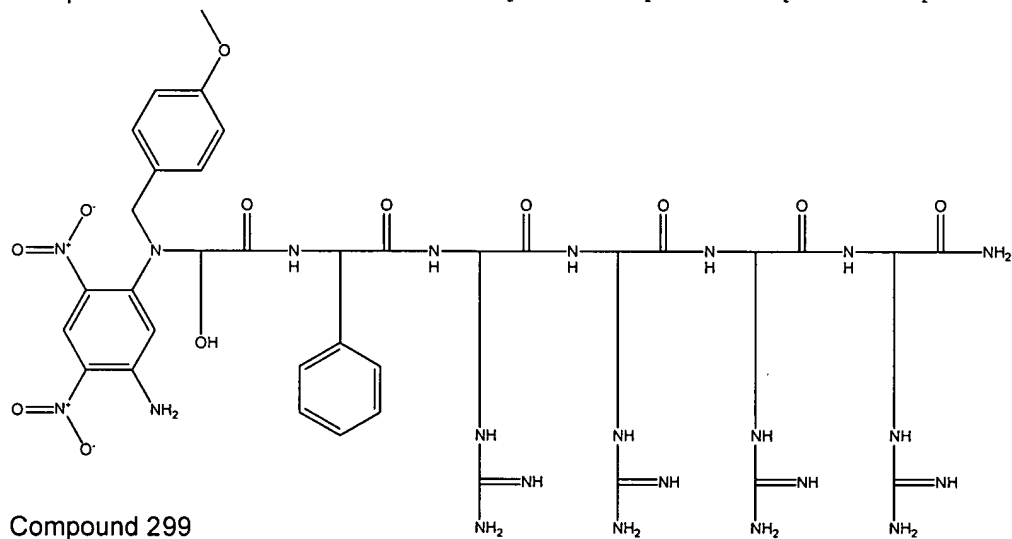


Compound 297

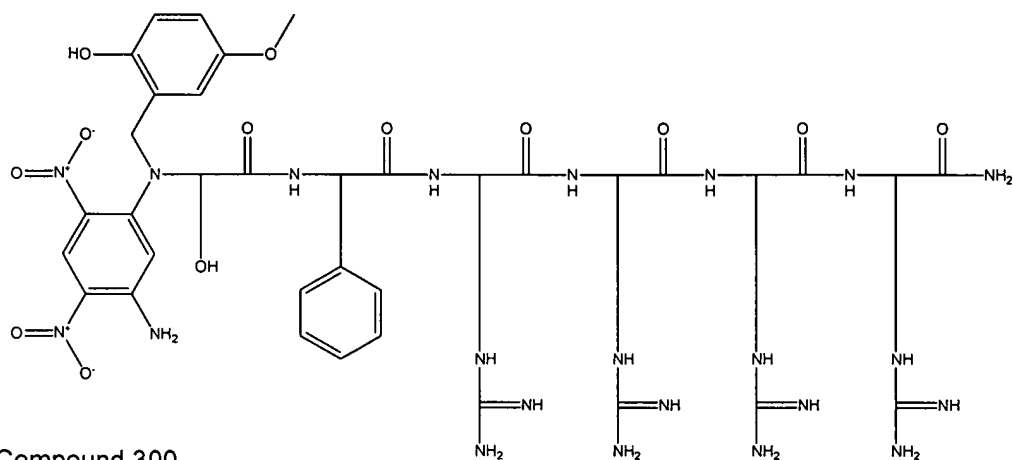
page 129 of 192



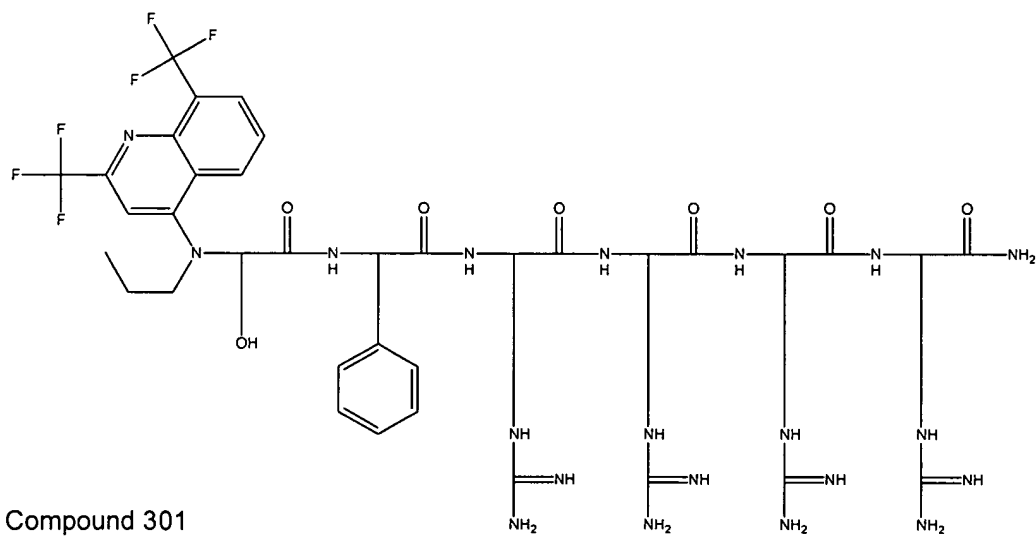
Compound 298



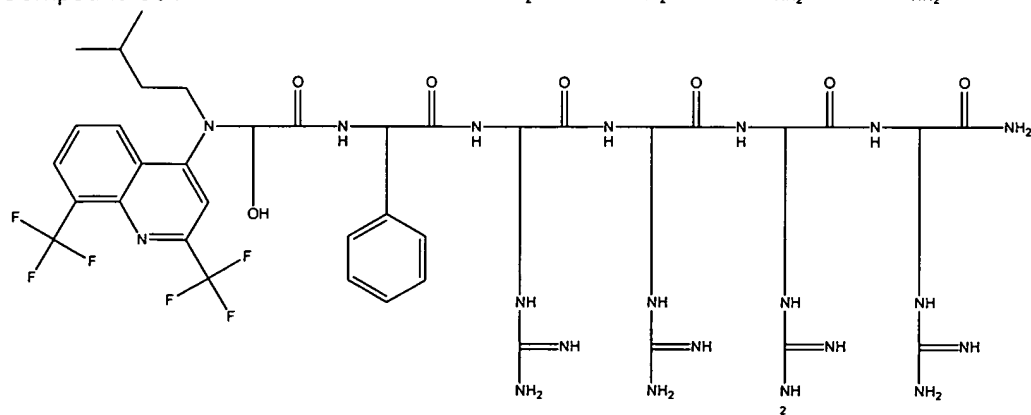
Compound 299



Compound 300



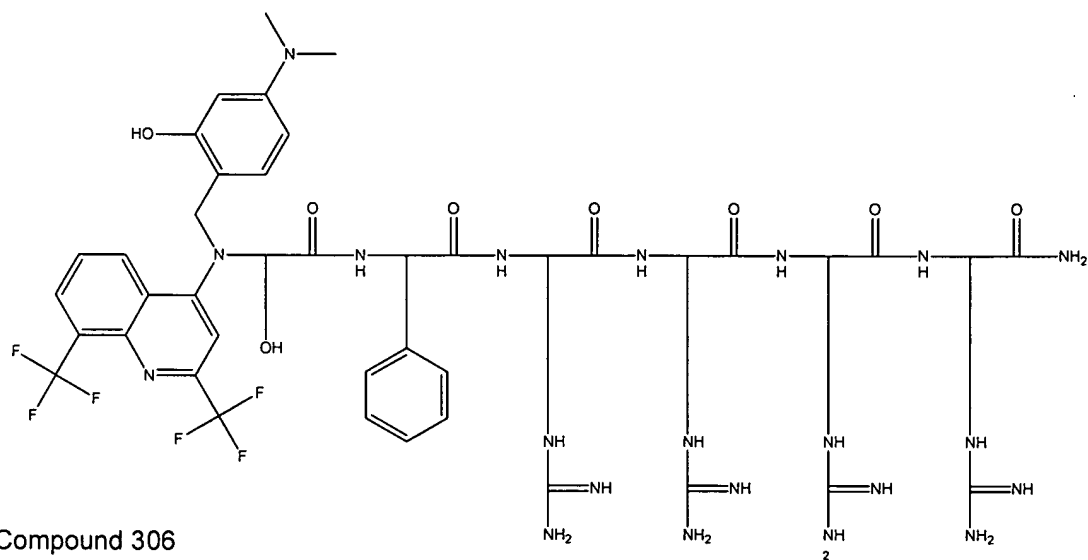
Compound 301



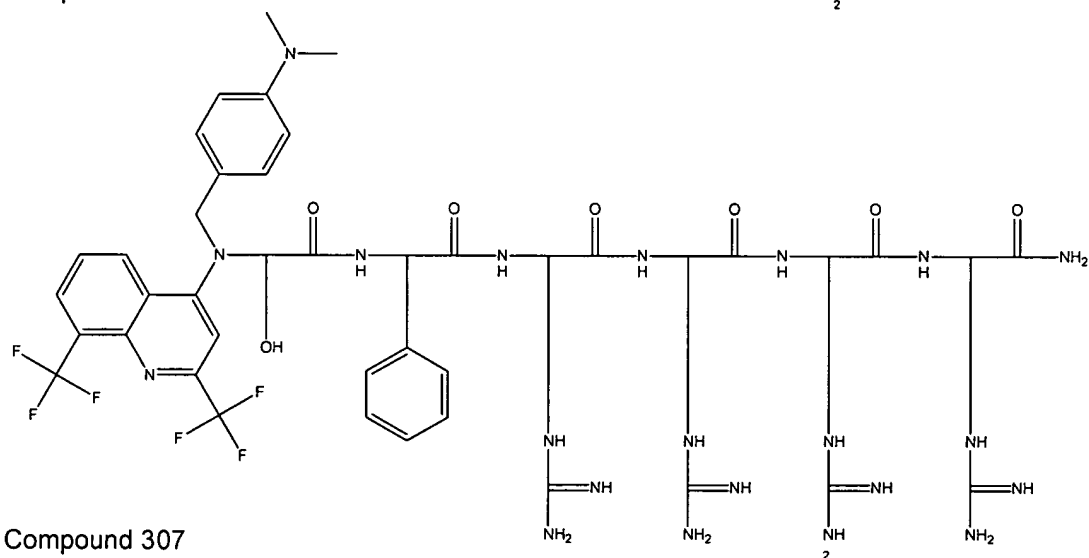
Compound 302

page 131 of 192



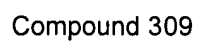


Compound 306

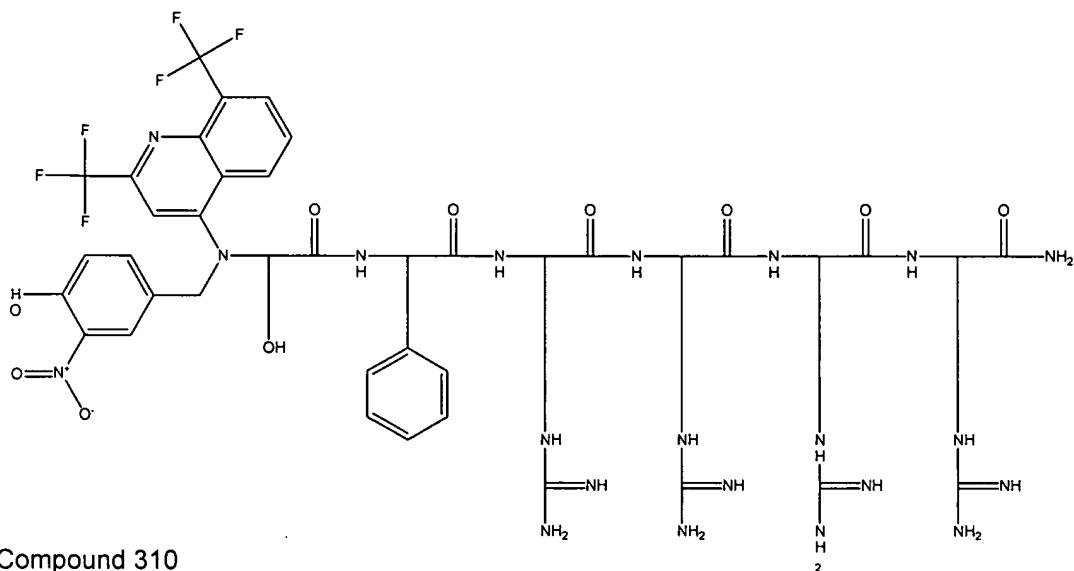


Compound 307

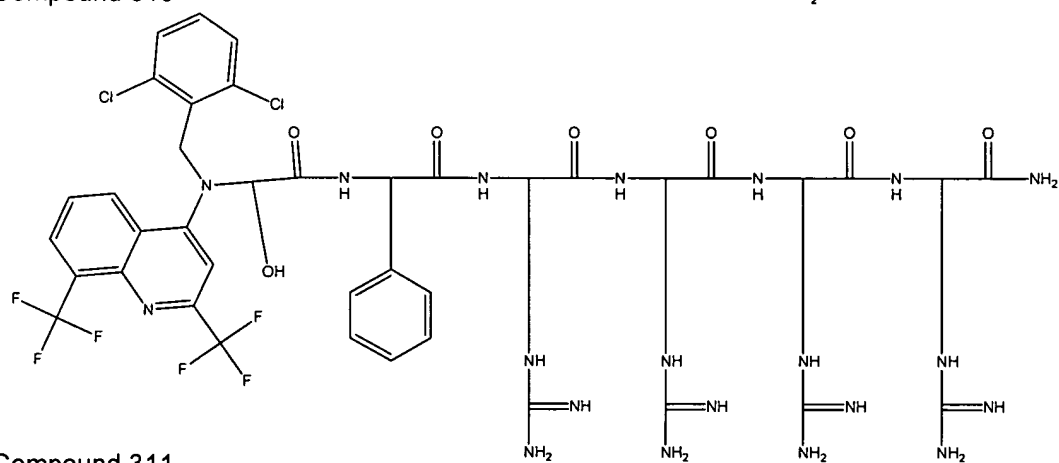
page 133 of 192



Applicant: David S. Lawrence
 Serial No.: 10/755,086
 Filed: January 9, 2004
 page 134 of 192



Compound 310



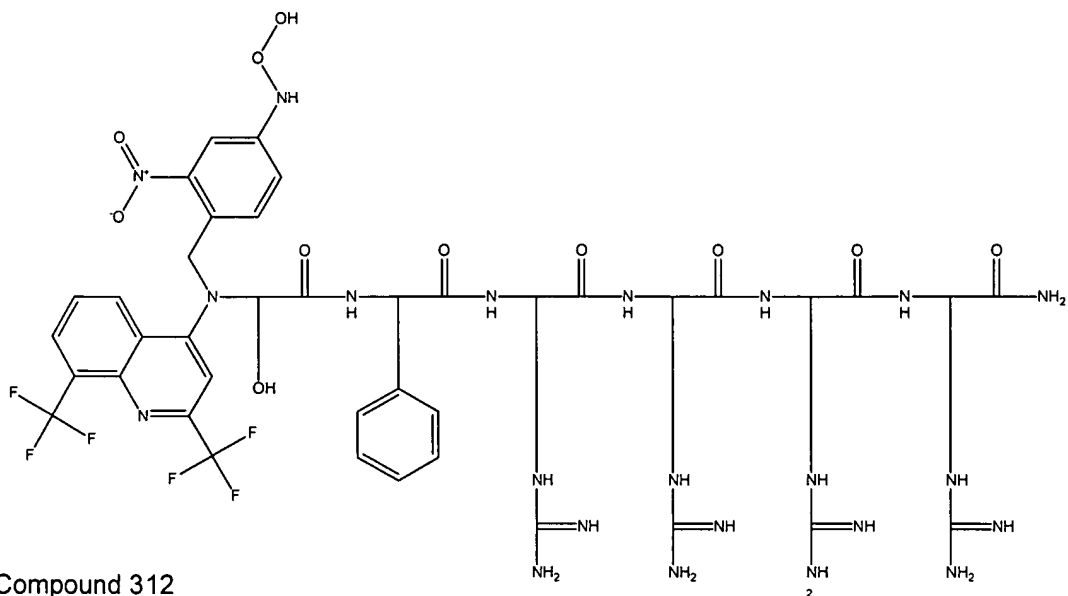
Compound 311

Applicant: David S. Lawrence

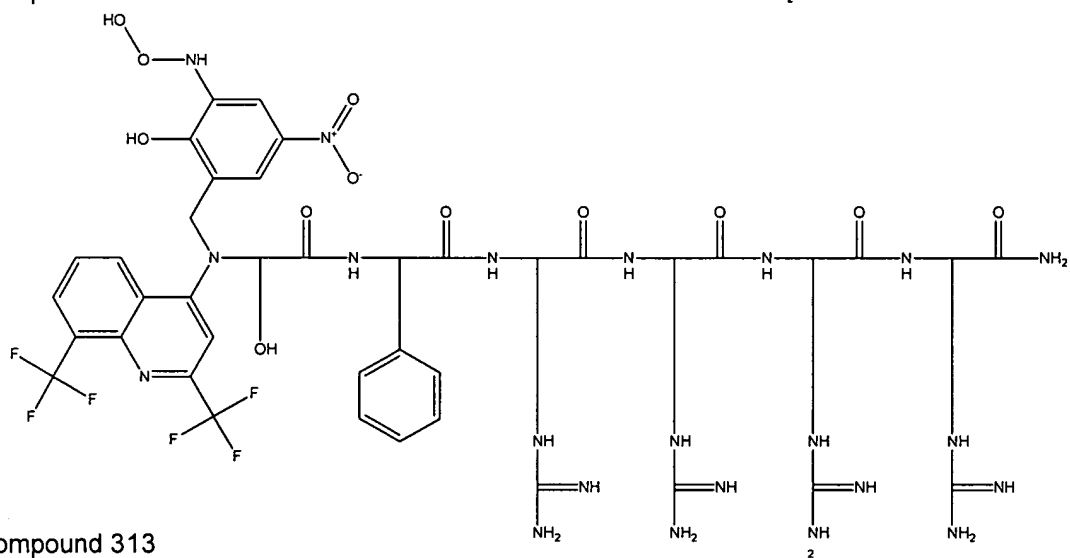
Serial No.: 10/755,086

Filed: January 9, 2004

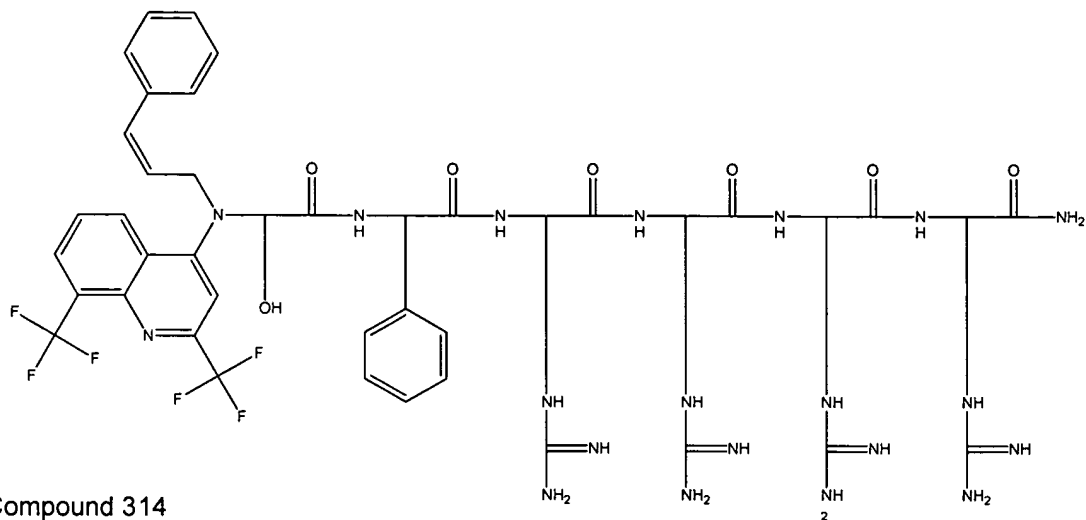
page 135 of 192



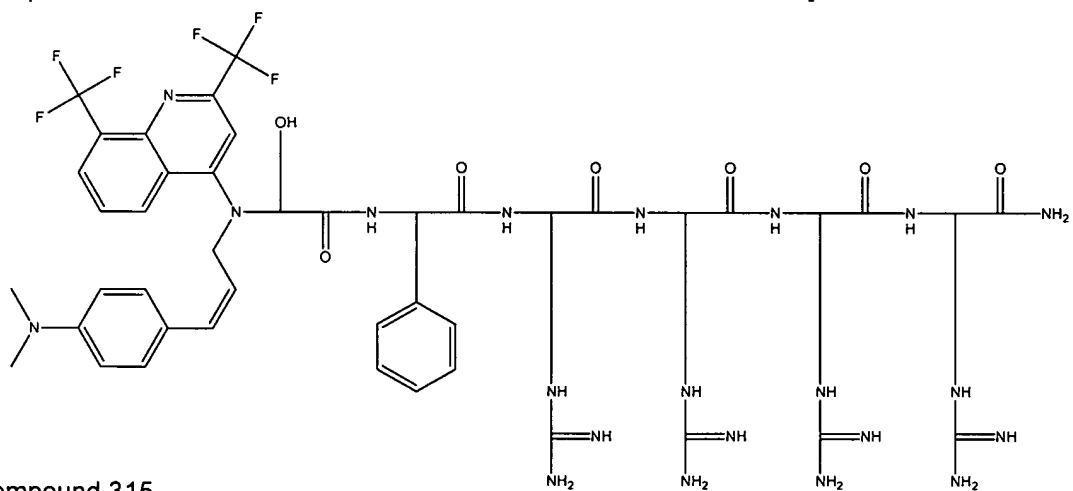
Compound 312



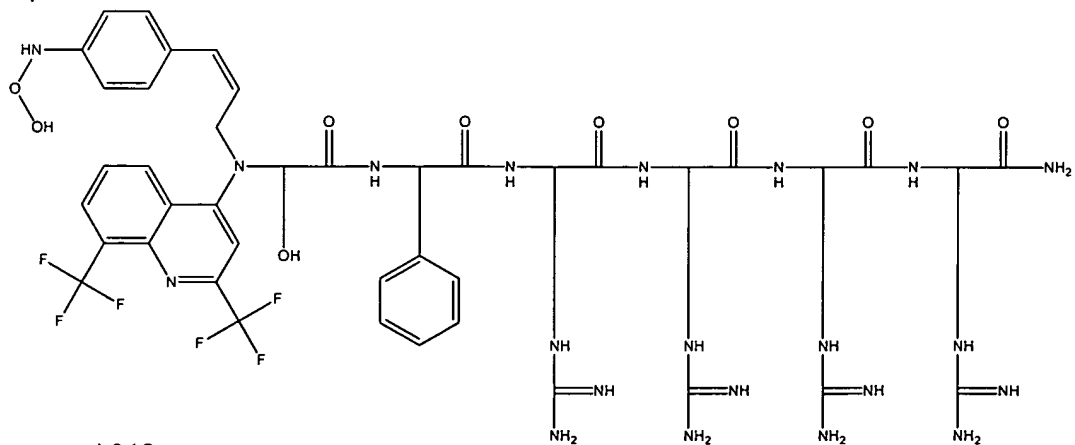
Compound 313



Compound 314

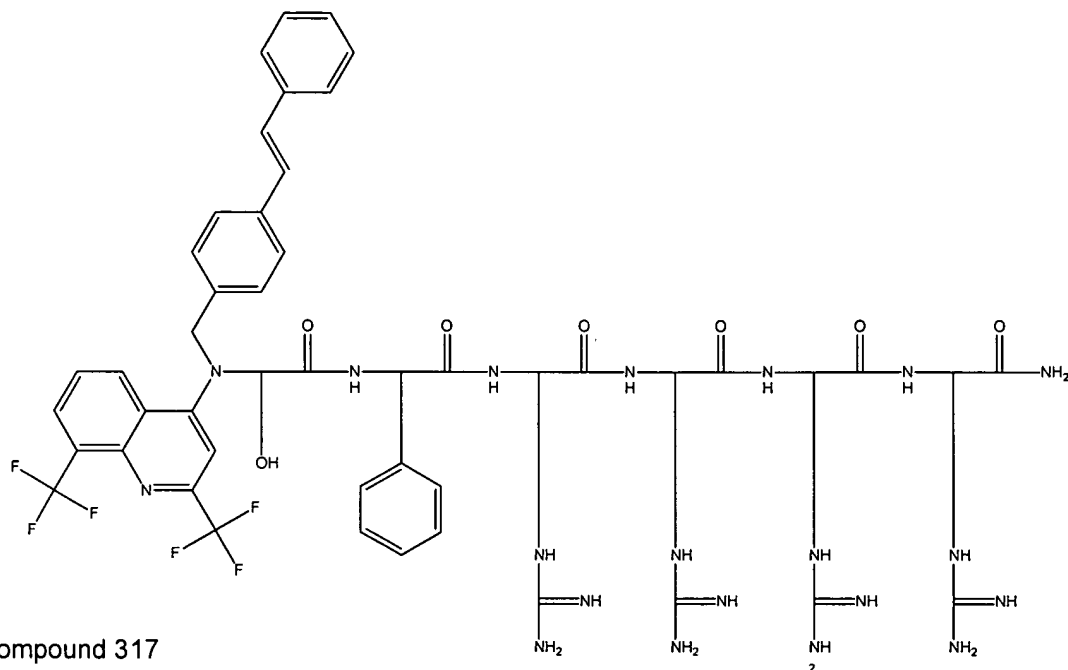


Compound 315

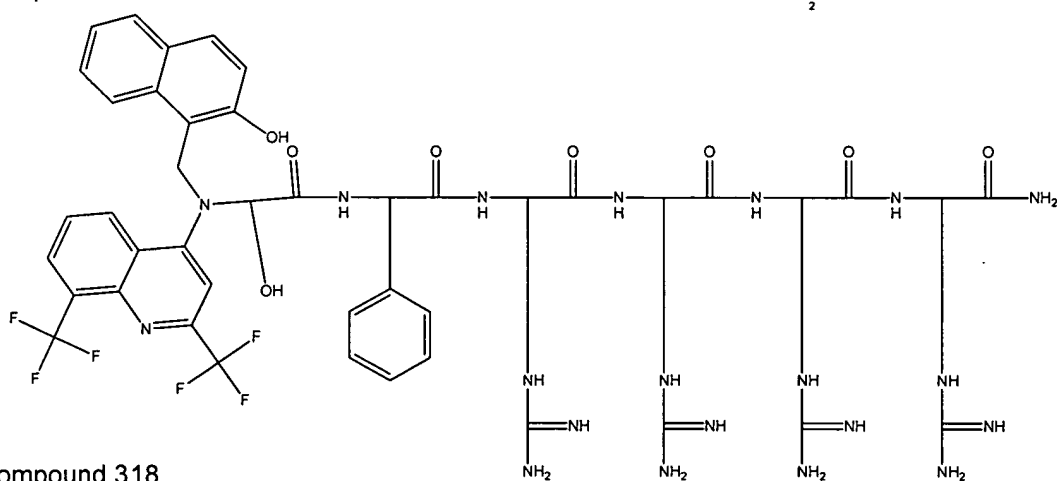


Compound 316

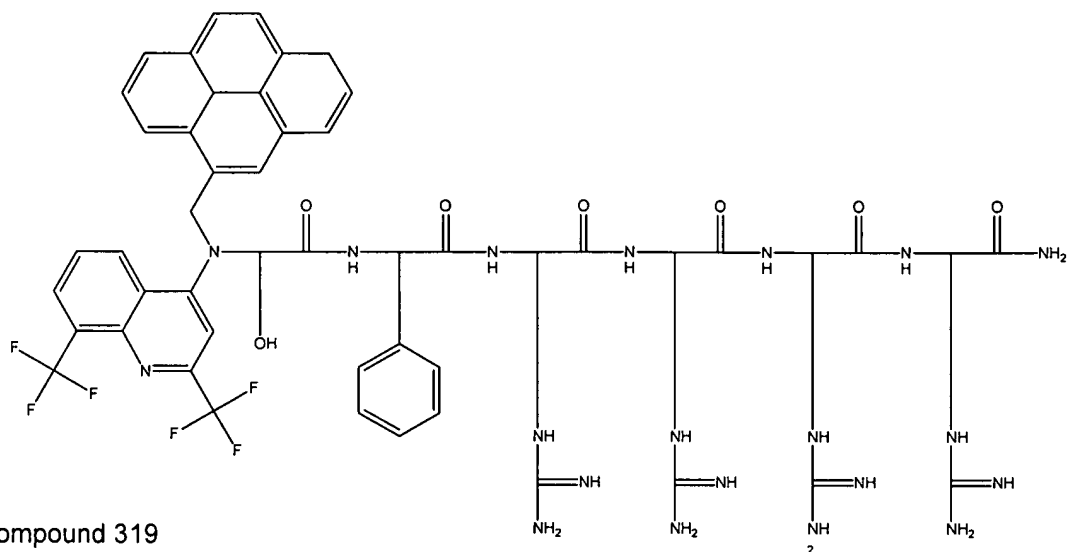
page 137 of 192



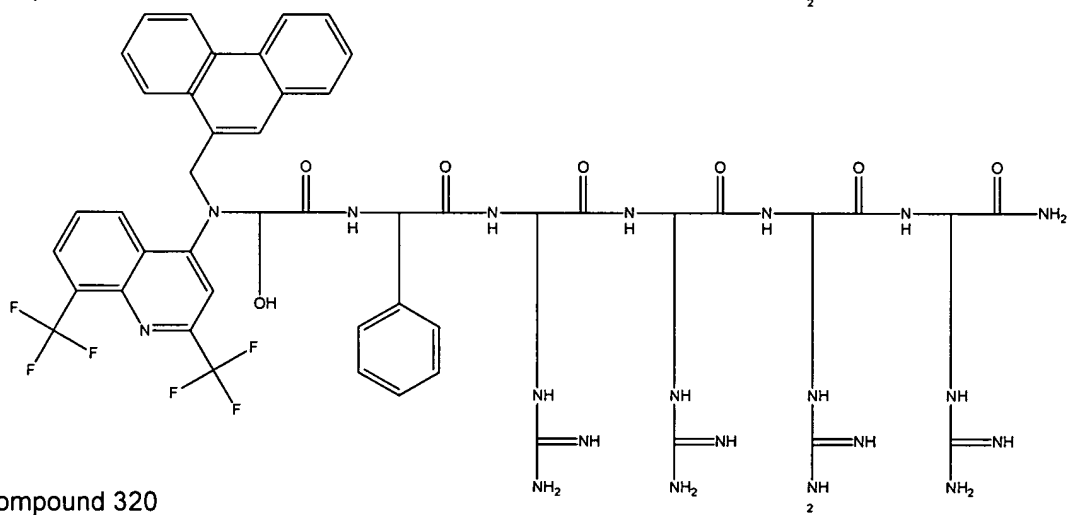
Compound 317



Compound 318

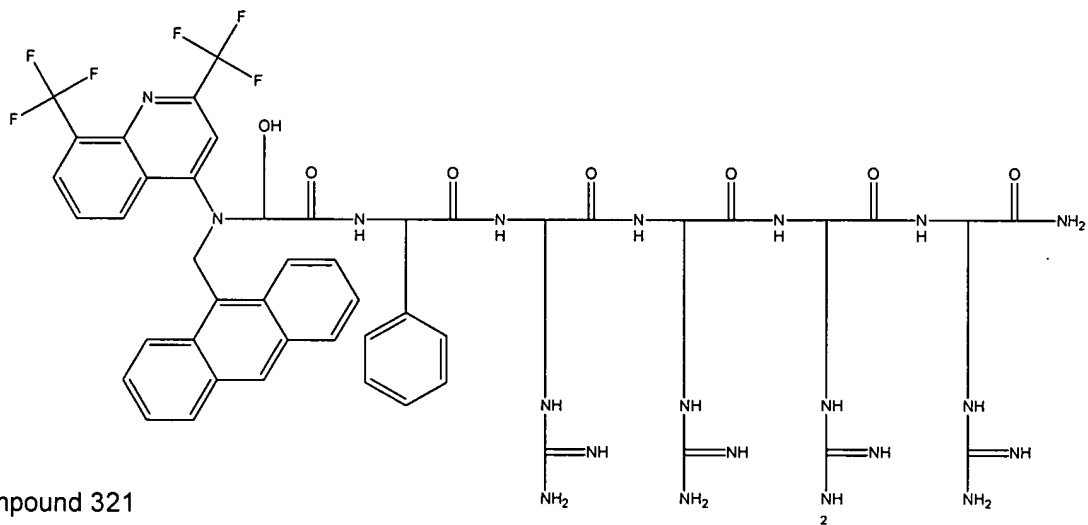


Compound 319

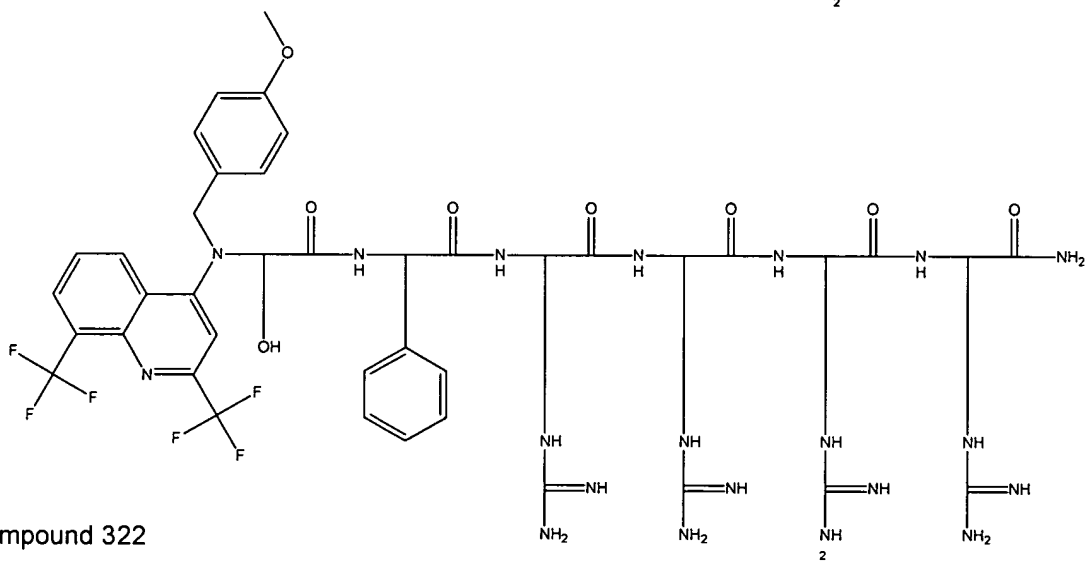


Compound 320

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 139 of 192

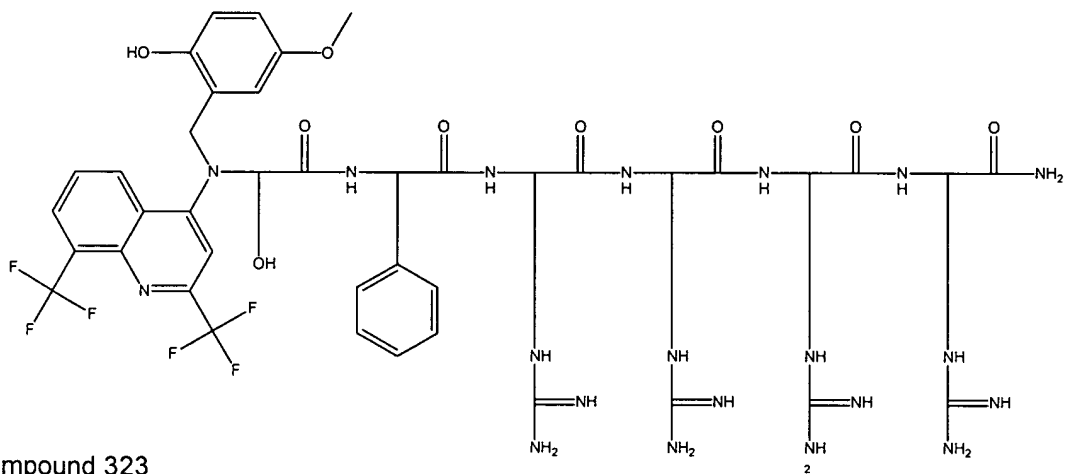


Compound 321

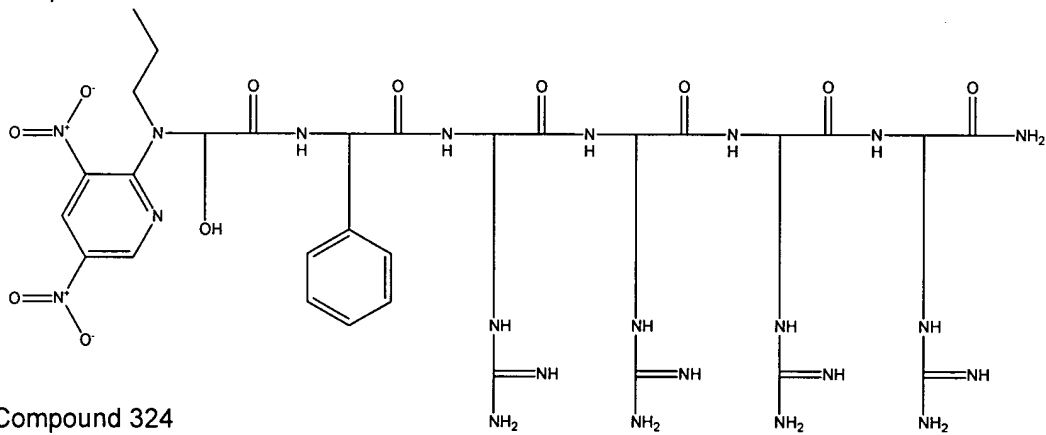


Compound 322

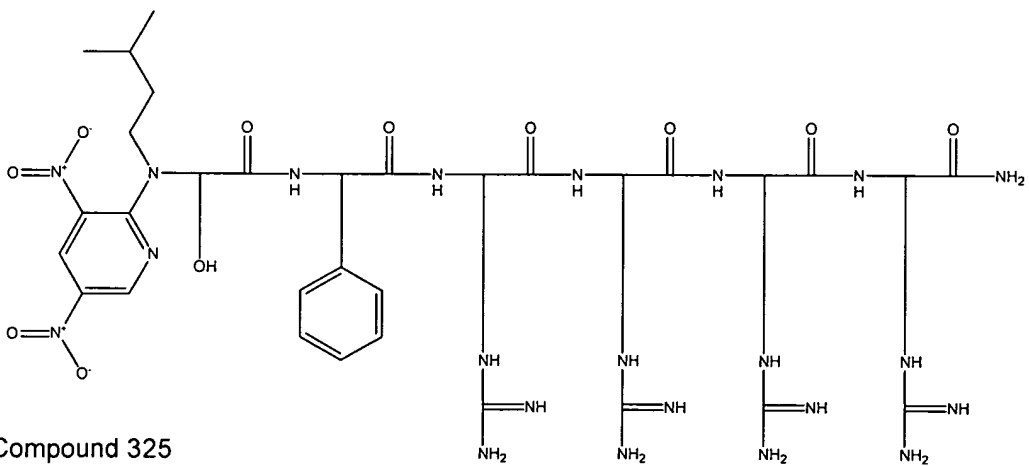
page 140 of 192



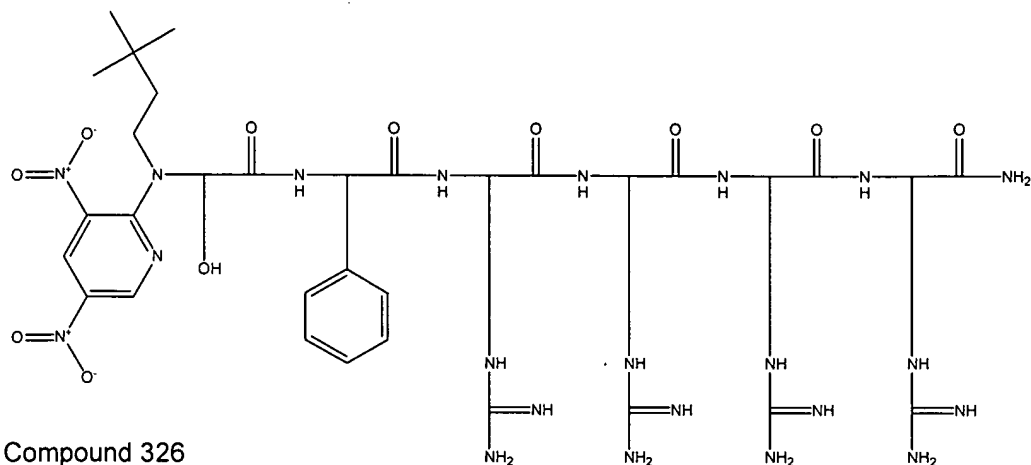
Compound 323



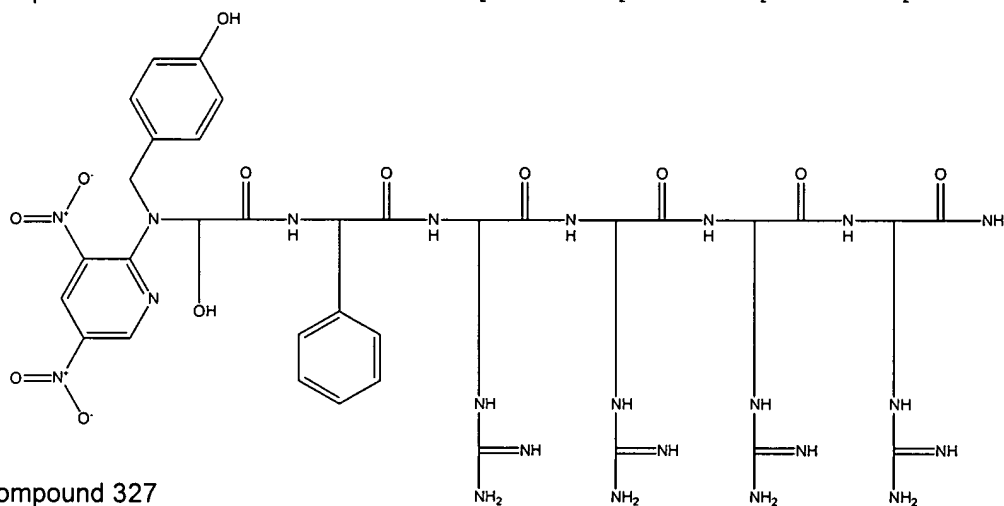
Compound 324



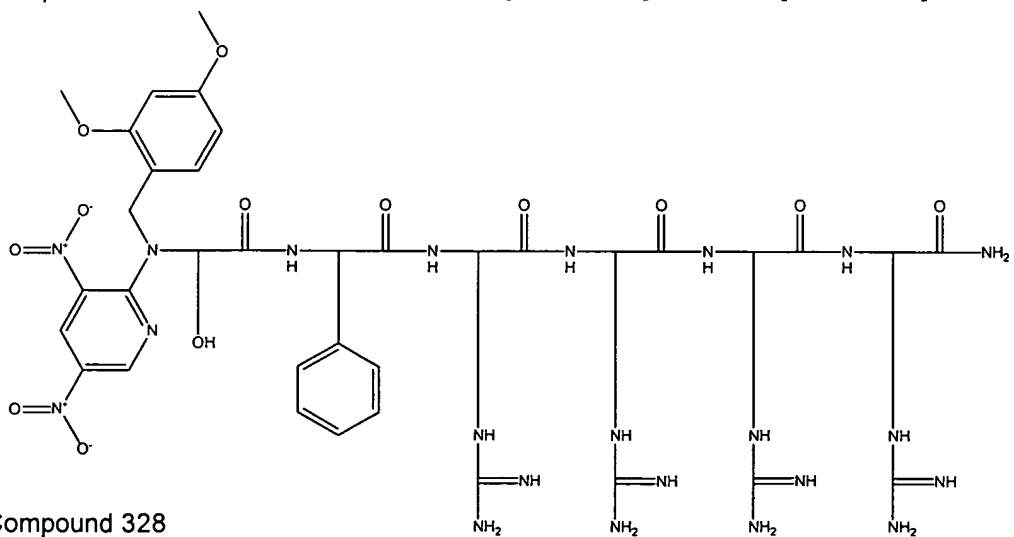
Compound 325



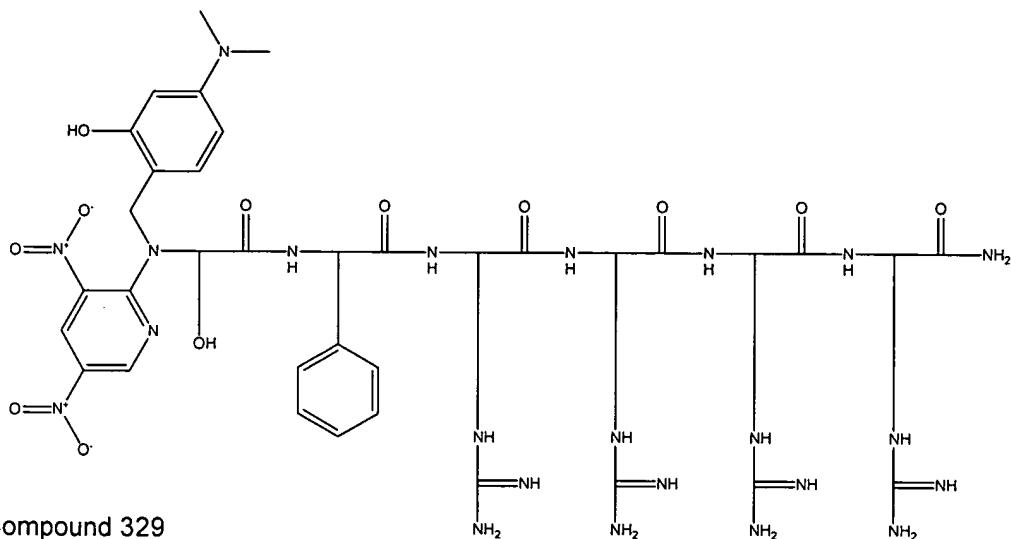
Compound 326



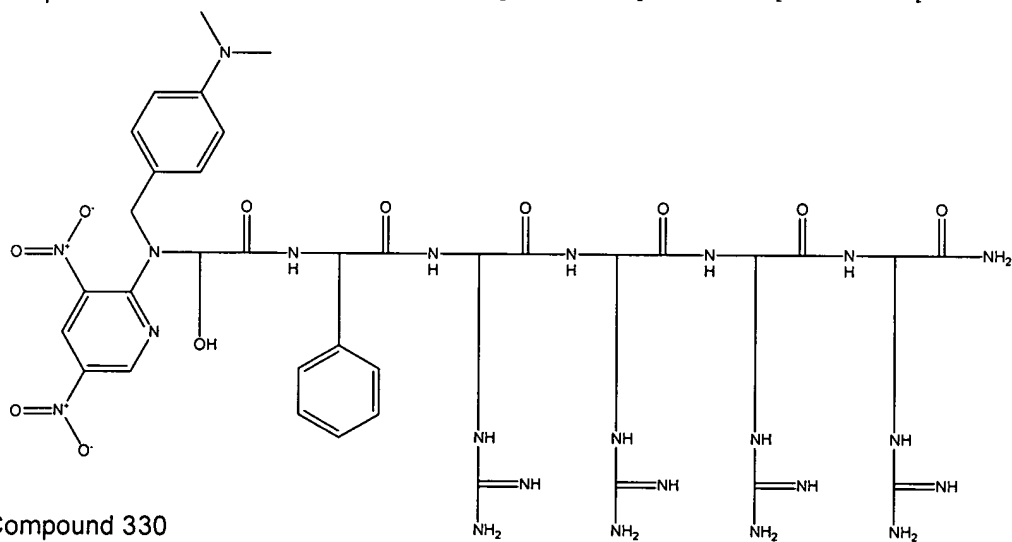
Compound 327



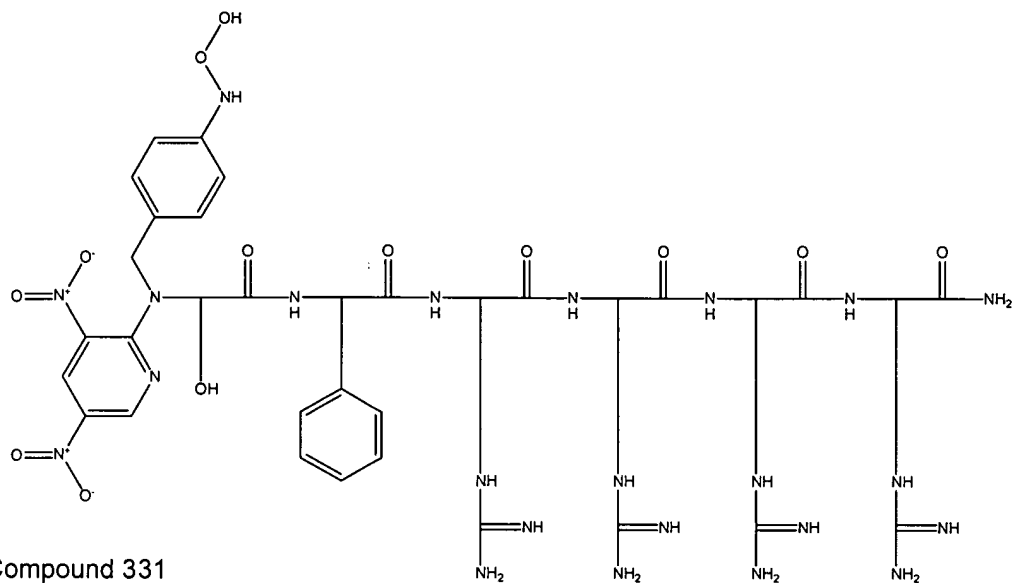
Compound 328



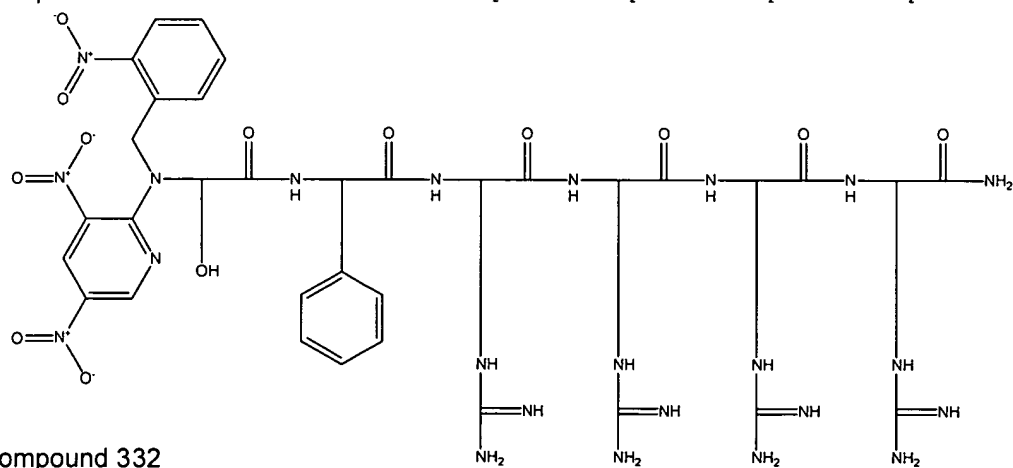
Compound 329



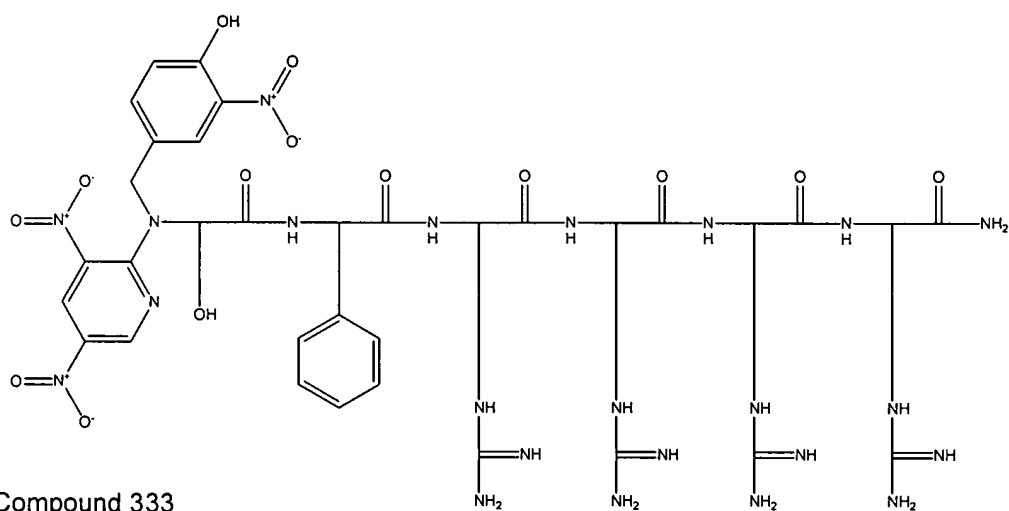
Compound 330



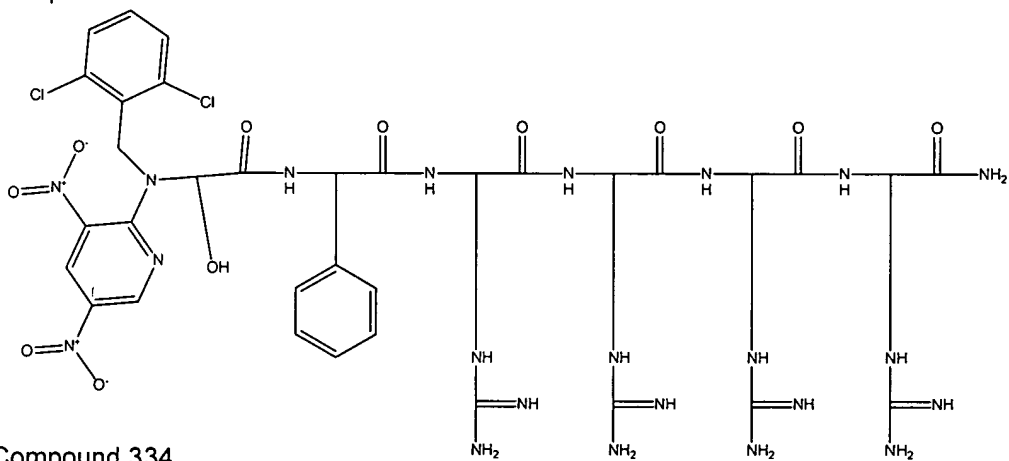
Compound 331



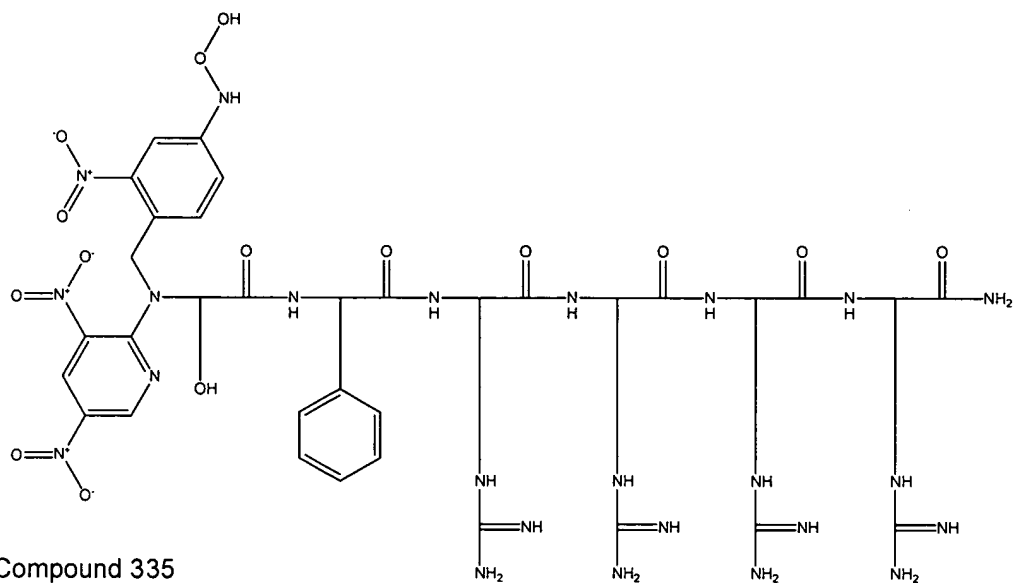
Compound 332



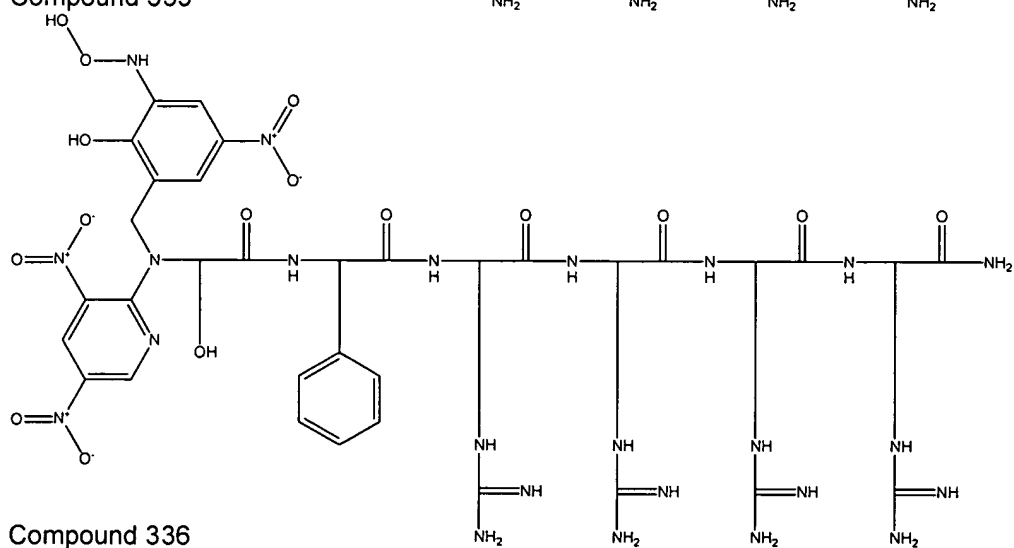
Compound 333



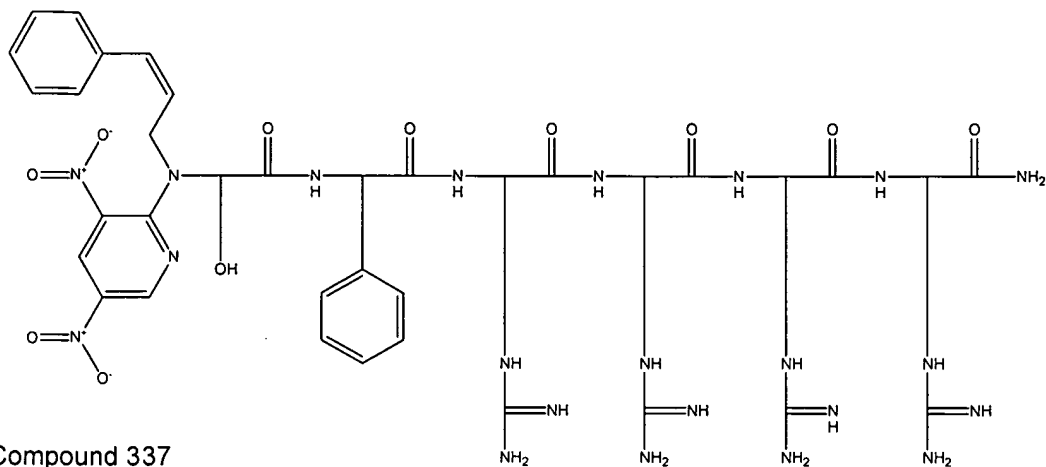
Compound 334



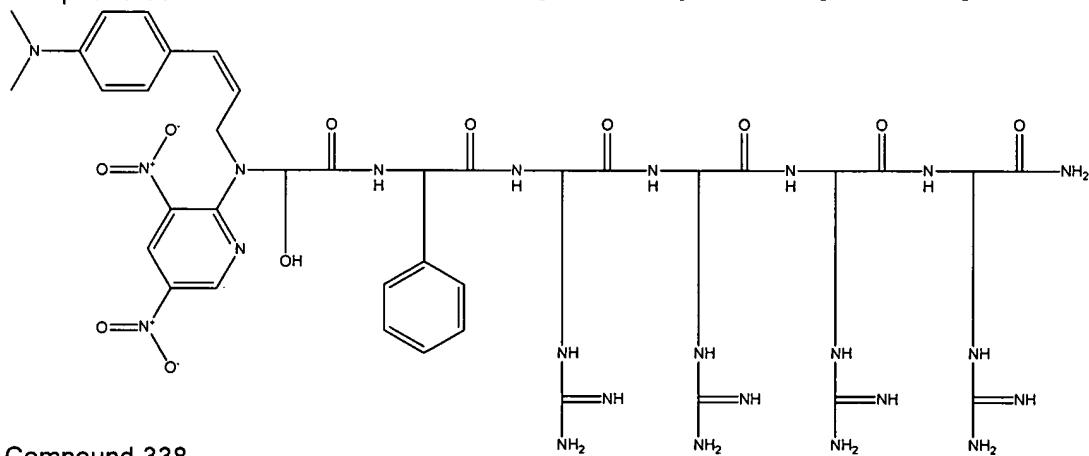
Compound 335



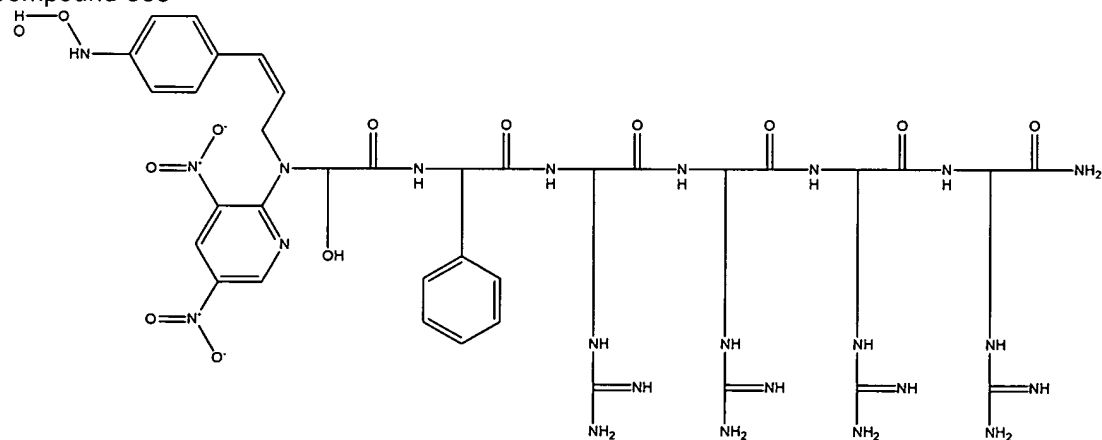
Compound 336



Compound 337

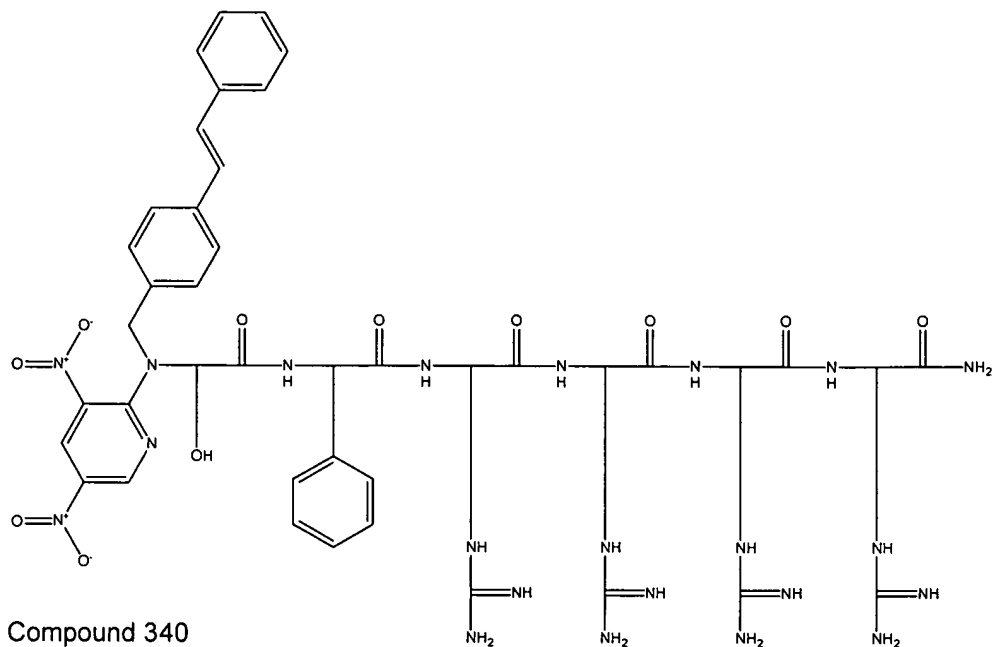


Compound 338

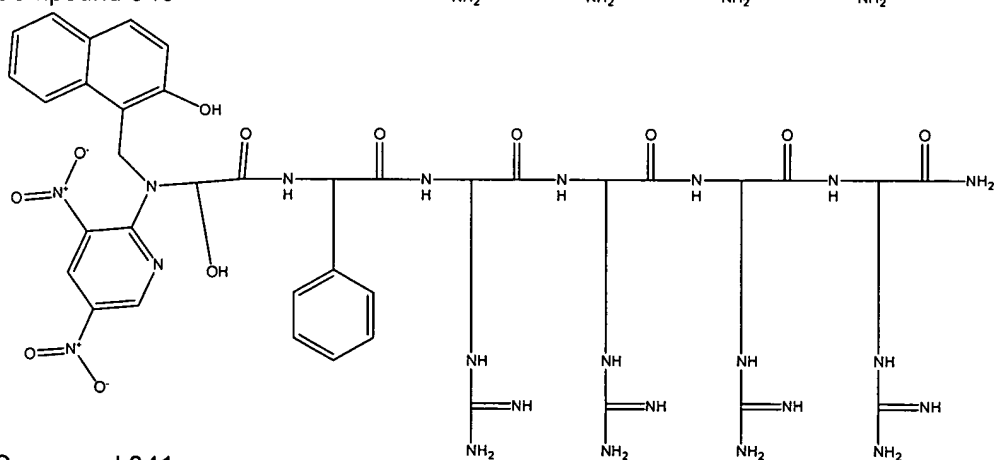


Compound 339

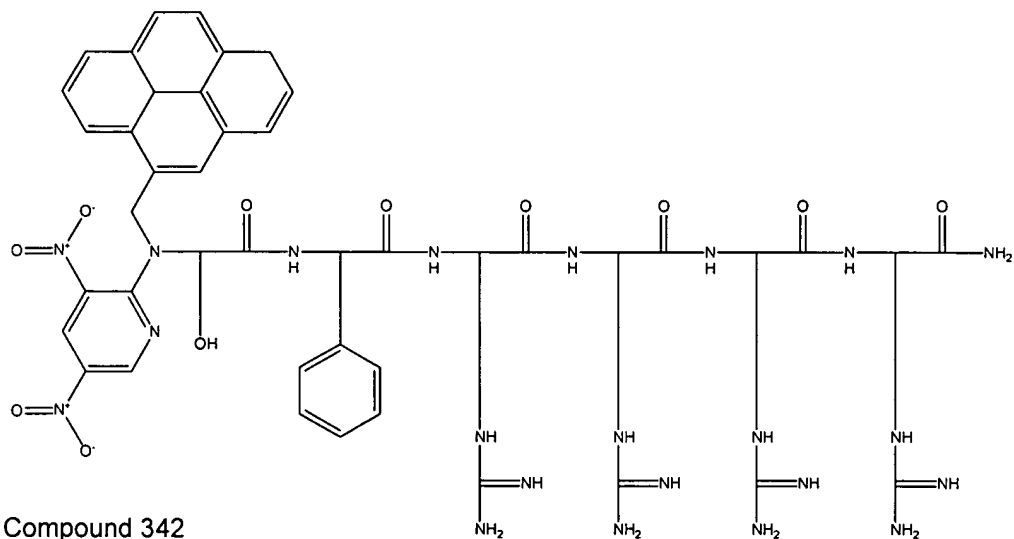
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 147 of 192



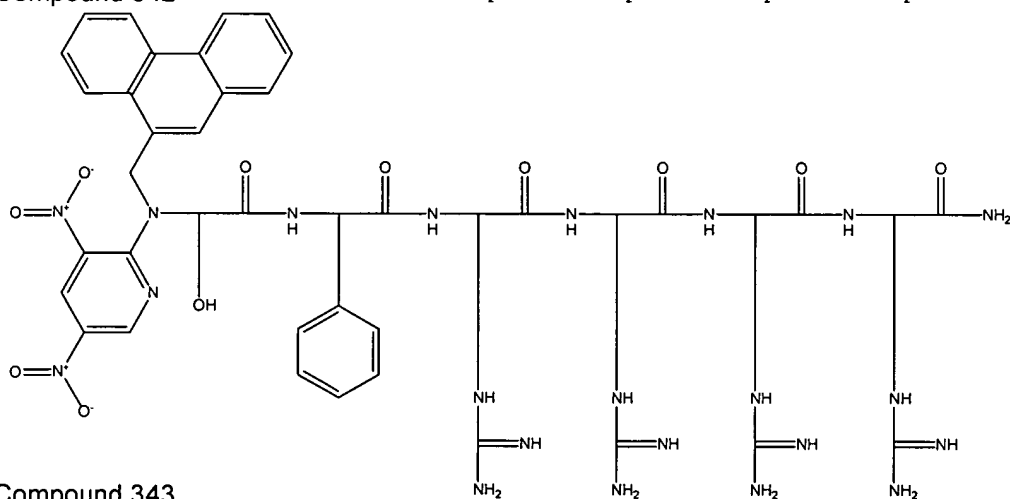
Compound 340



Compound 341

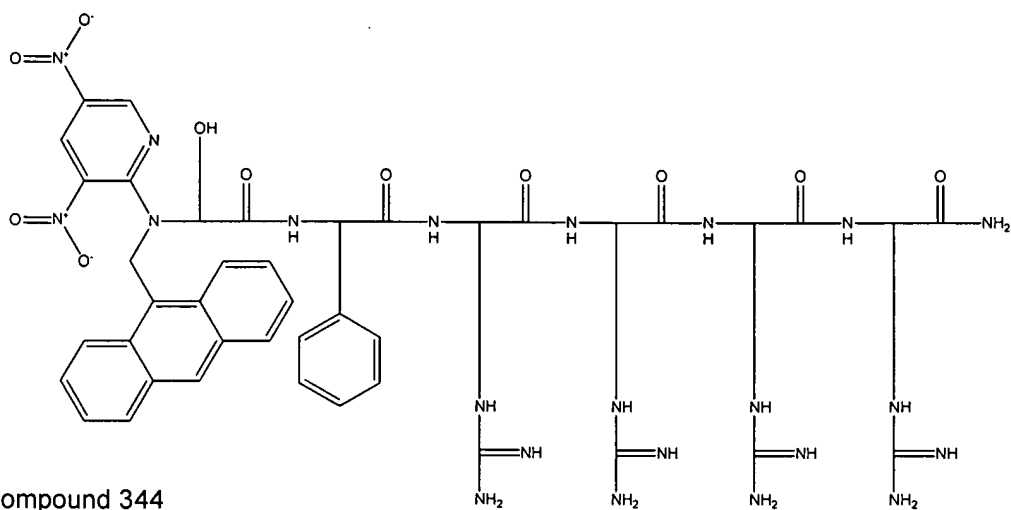


Compound 342

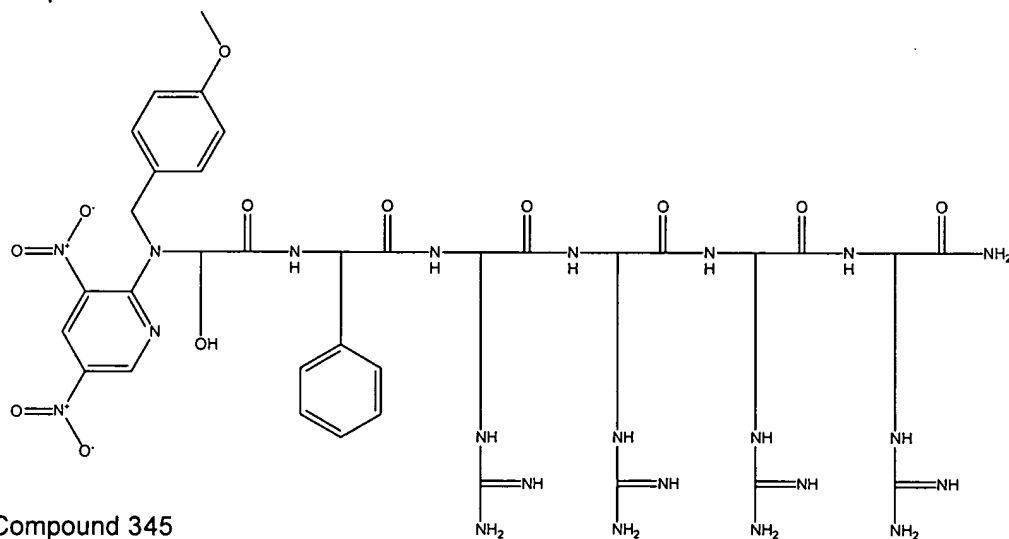


Compound 343

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 149 of 192



Compound 344



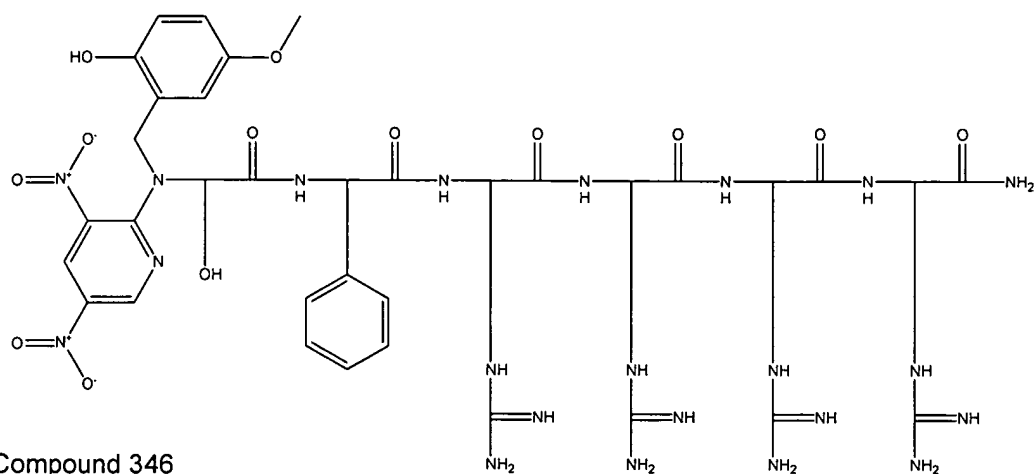
Compound 345

Applicant: David S. Lawrence

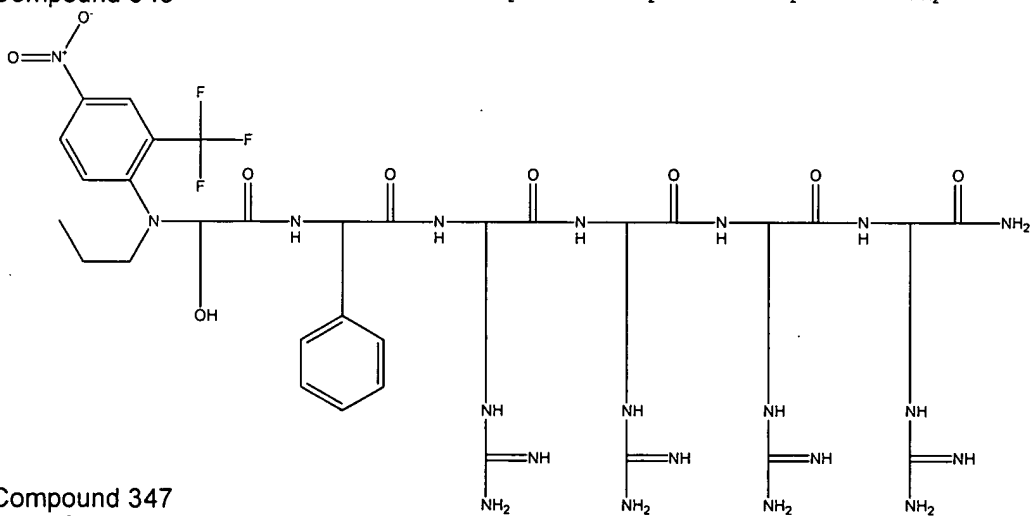
Serial No.: 10/755,086

Filed: January 9, 2004

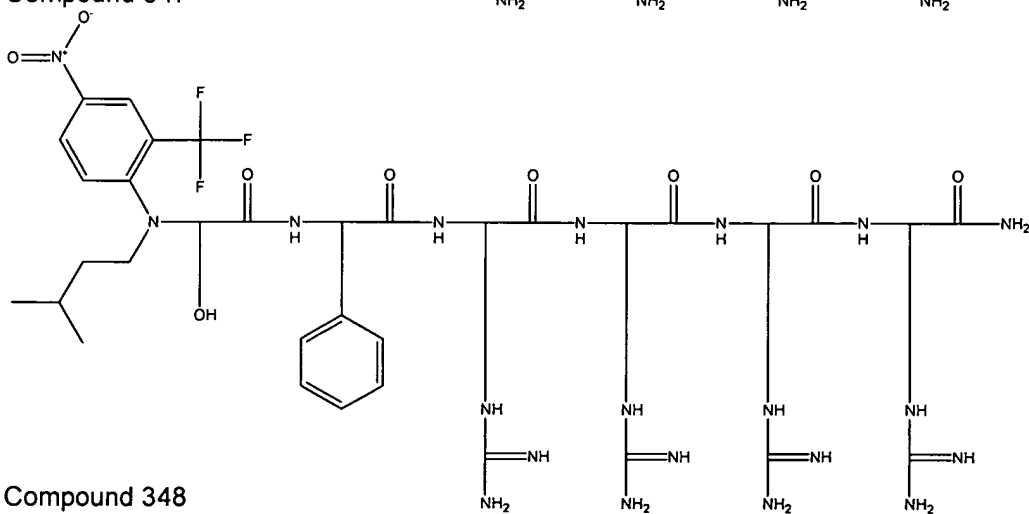
page 150 of 192



Compound 346

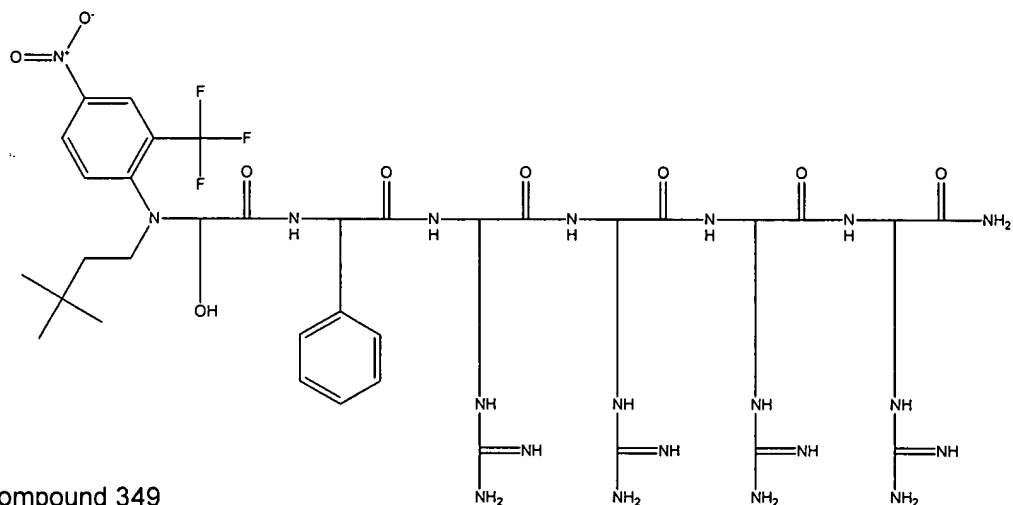


Compound 347

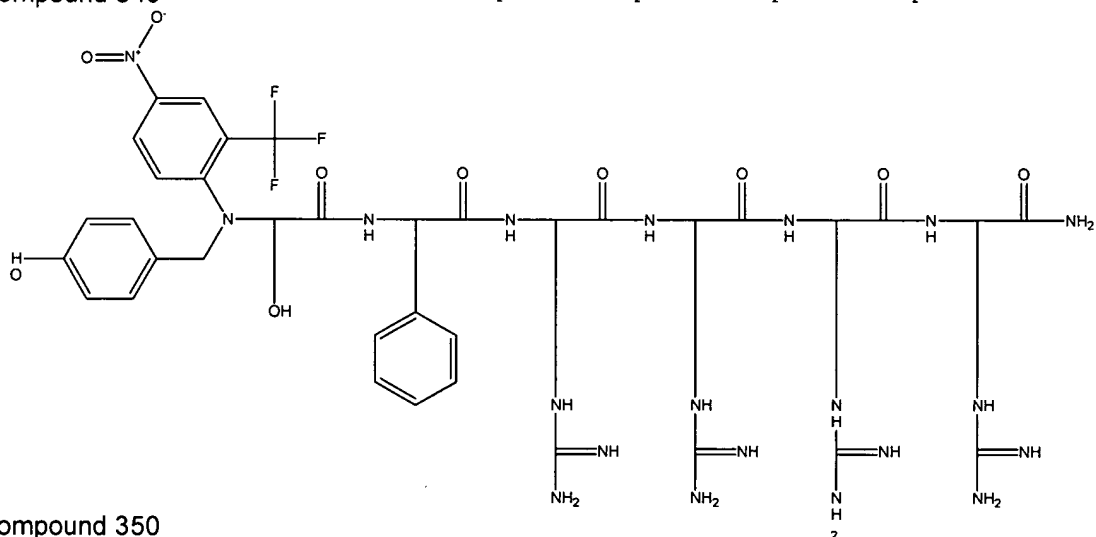


Compound 348

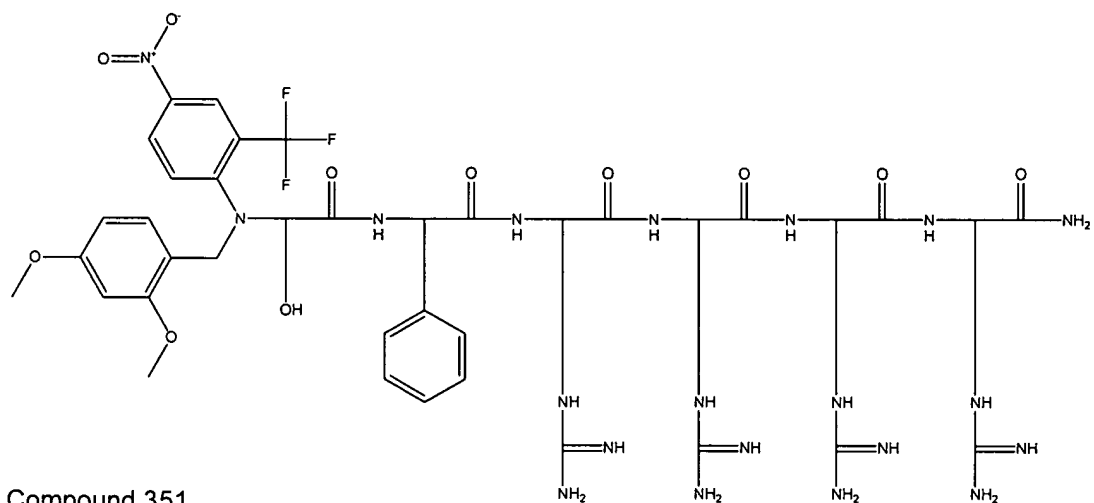
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 151 of 192



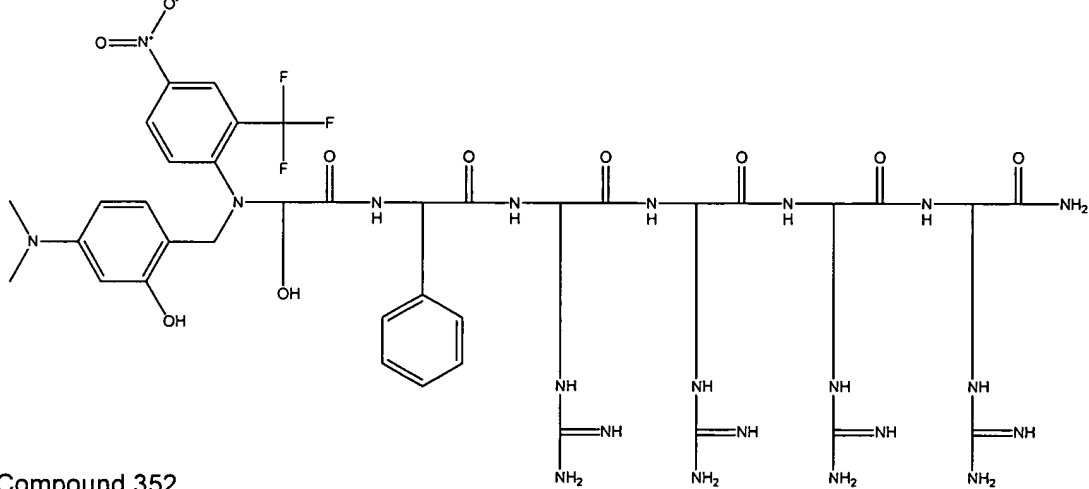
Compound 349



Compound 350

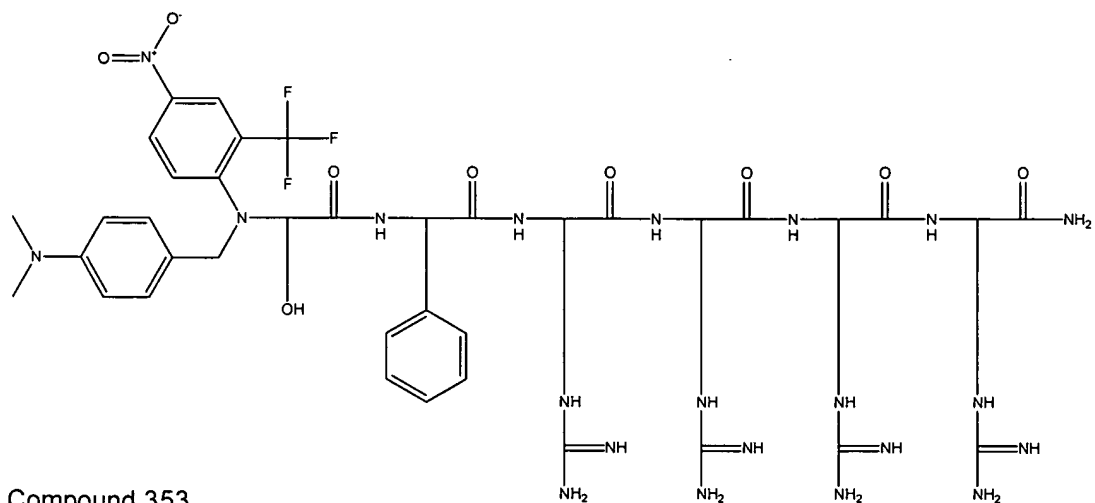


Compound 351

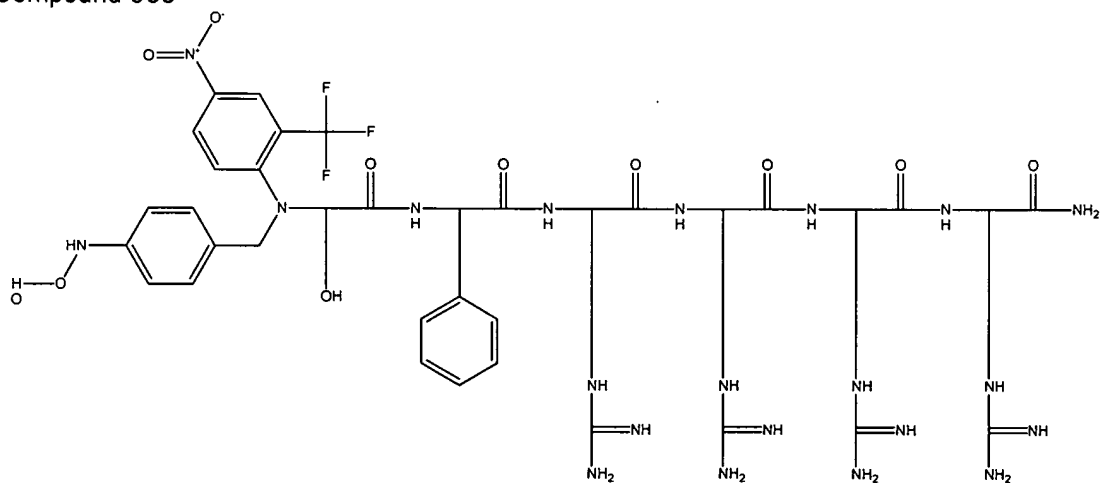


Compound 352

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 153 of 192

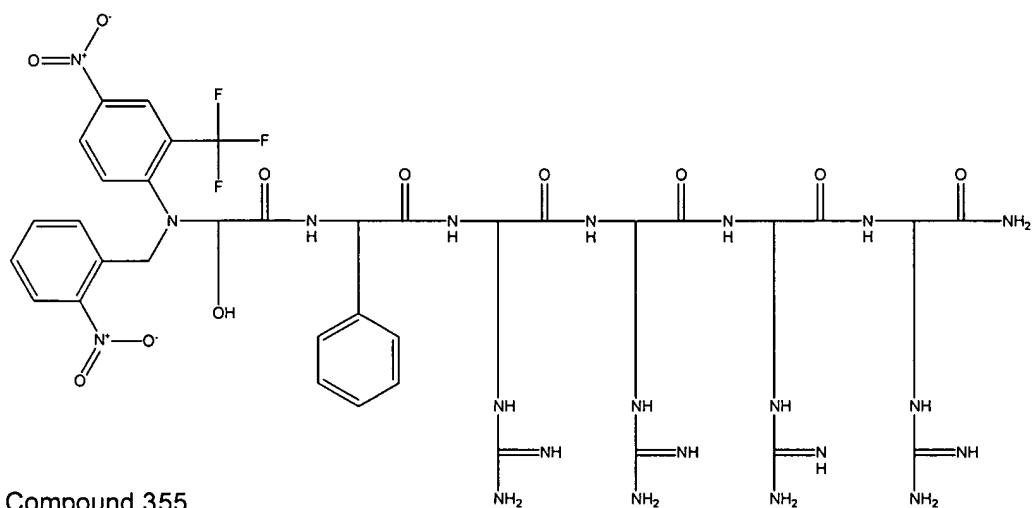


Compound 353

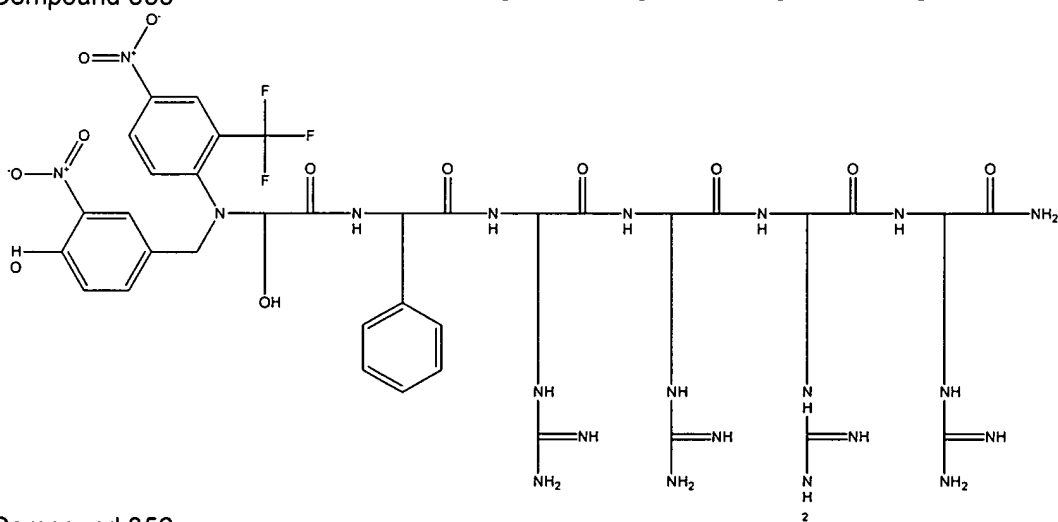


Compound 354

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 154 of 192

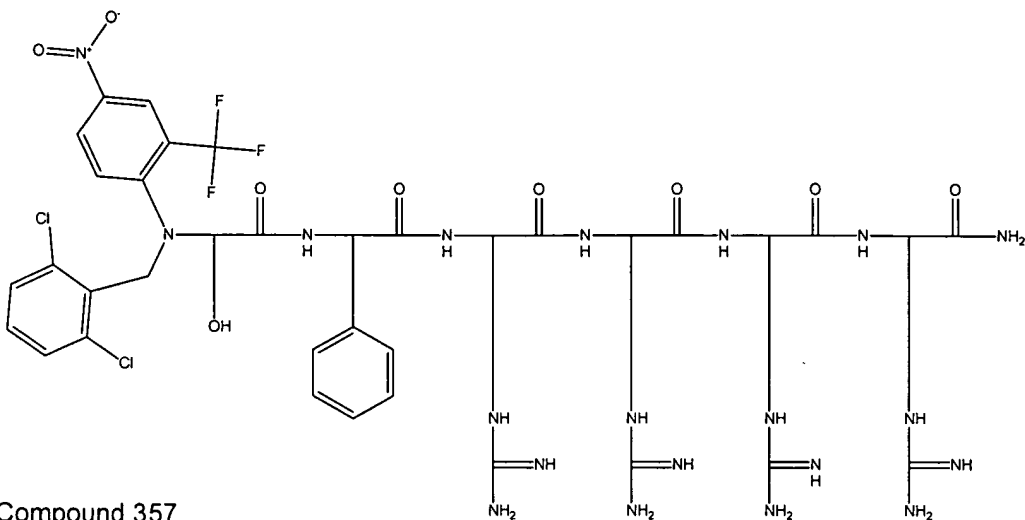


Compound 355

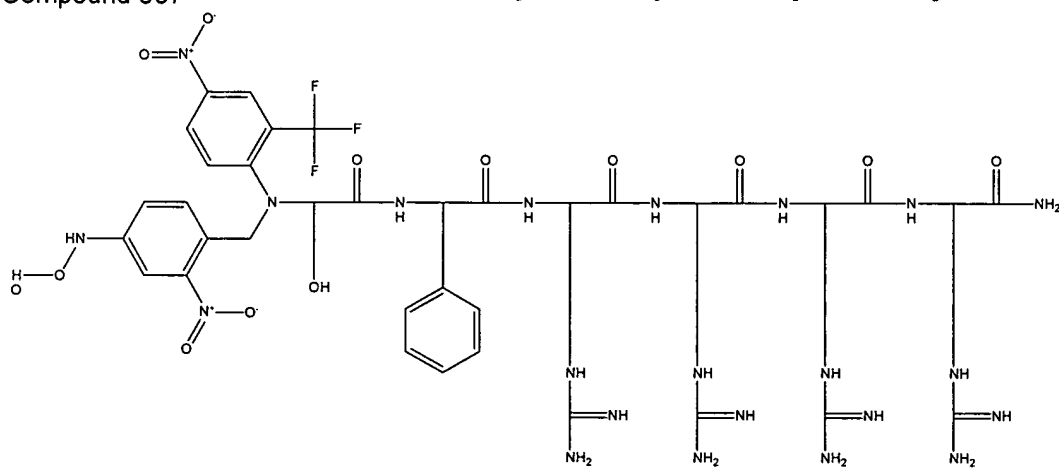


Compound 356

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 155 of 192

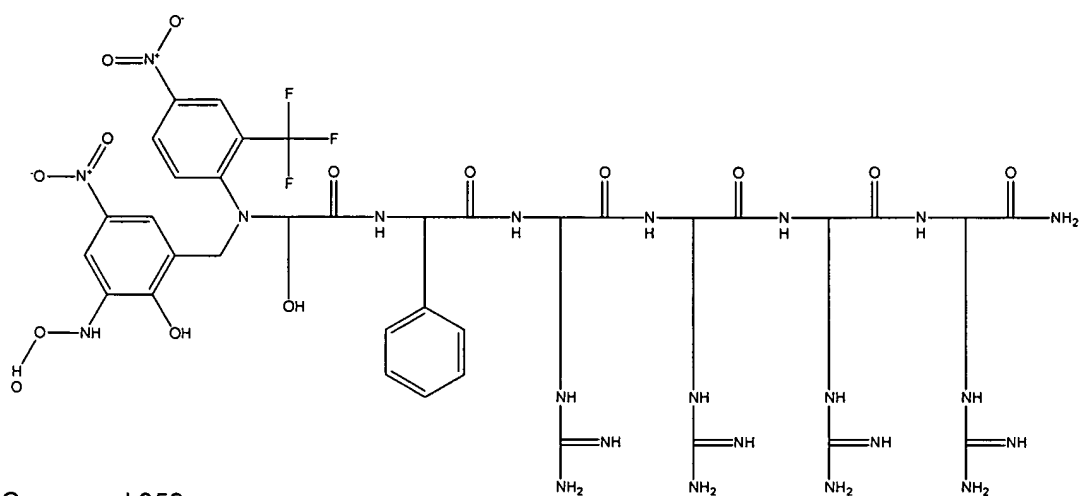


Compound 357

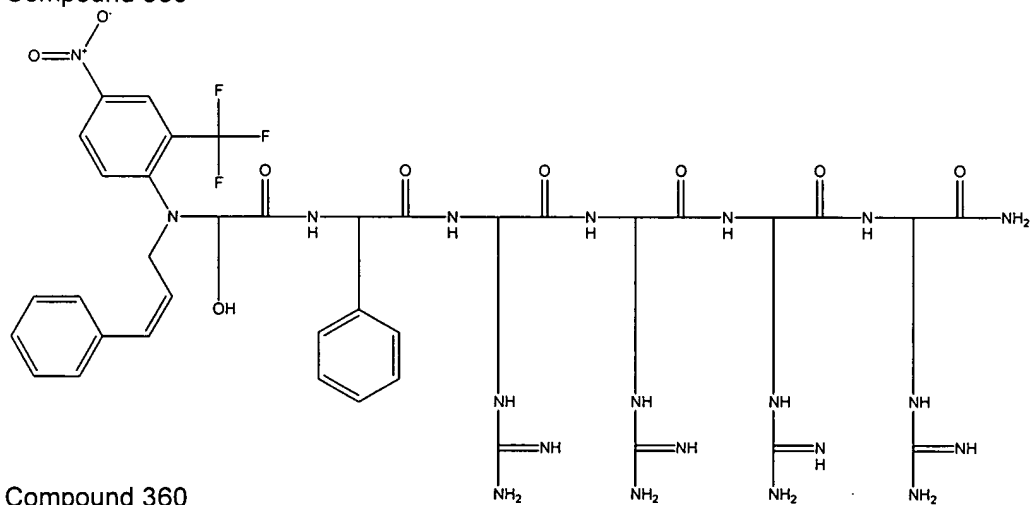


Compound 358

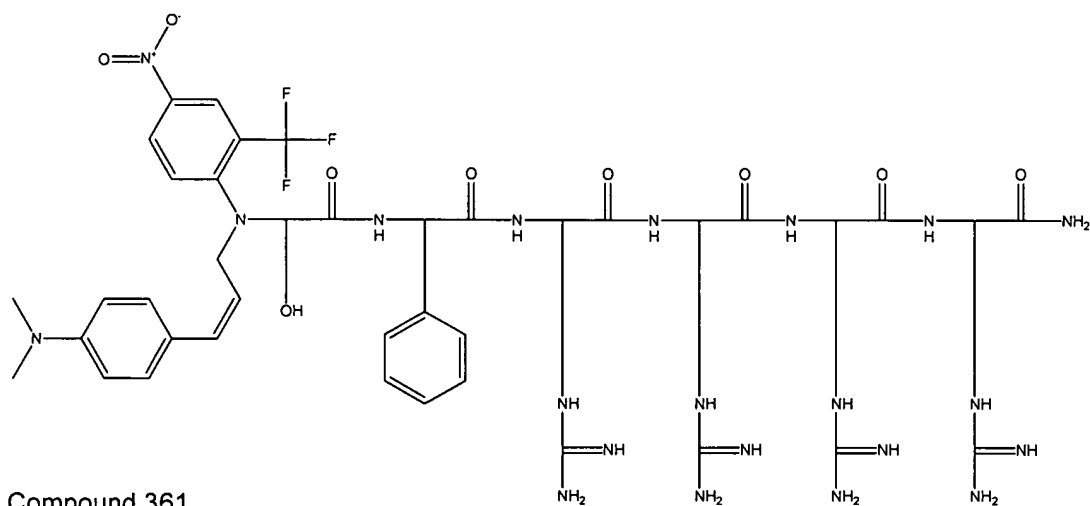
page 156 of 192



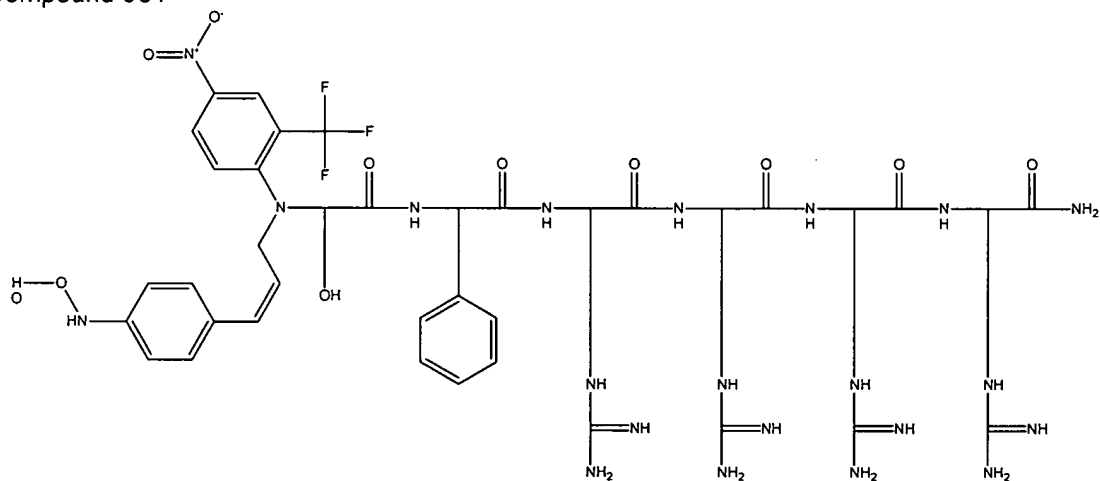
Compound 359



Compound 360

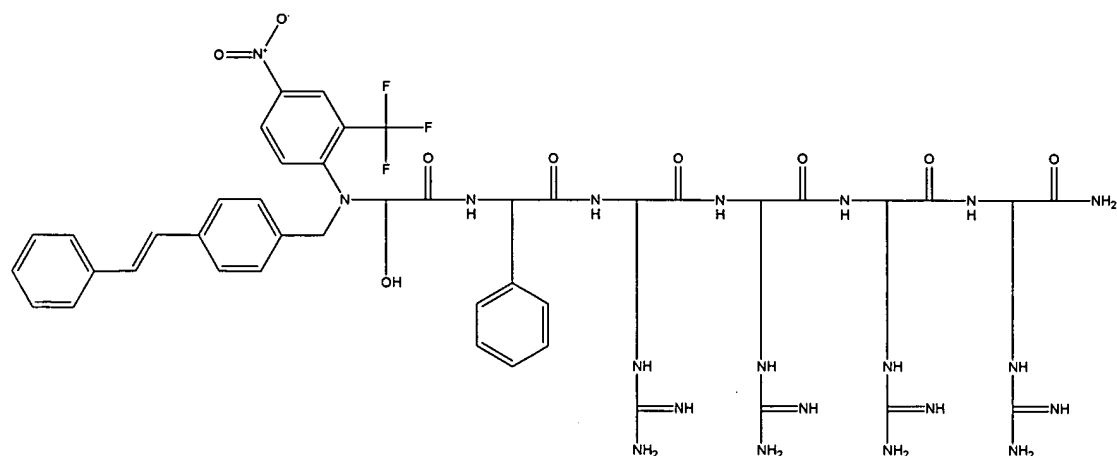


Compound 361

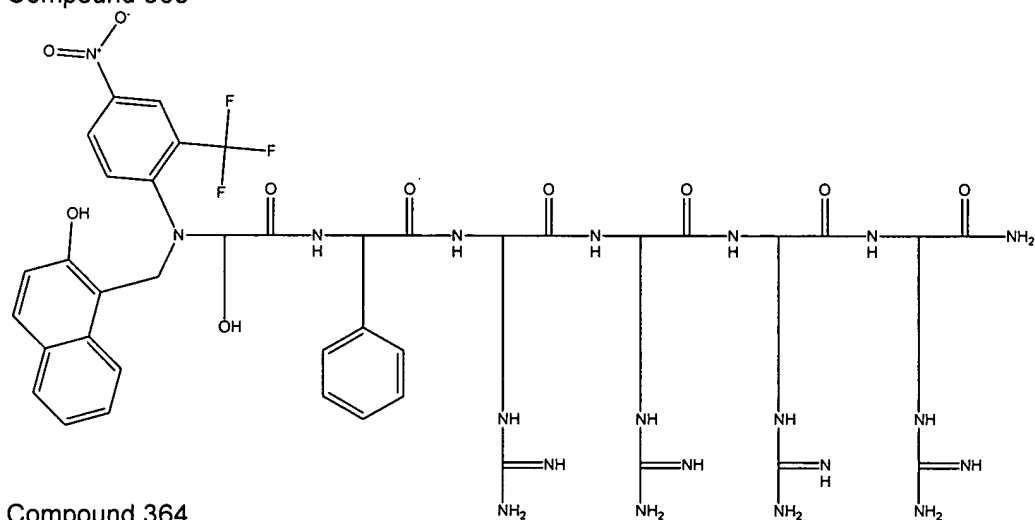


Compound 362

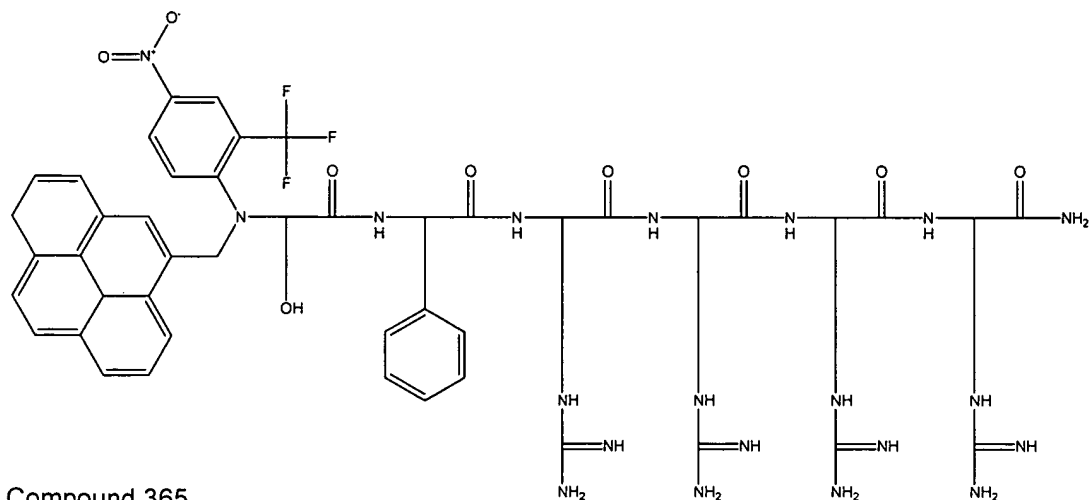
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 158 of 192



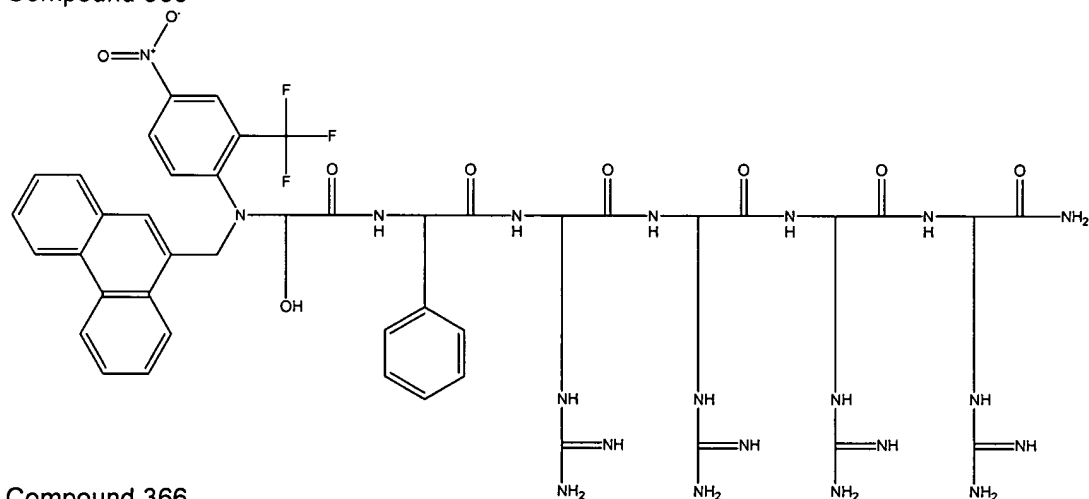
Compound 363



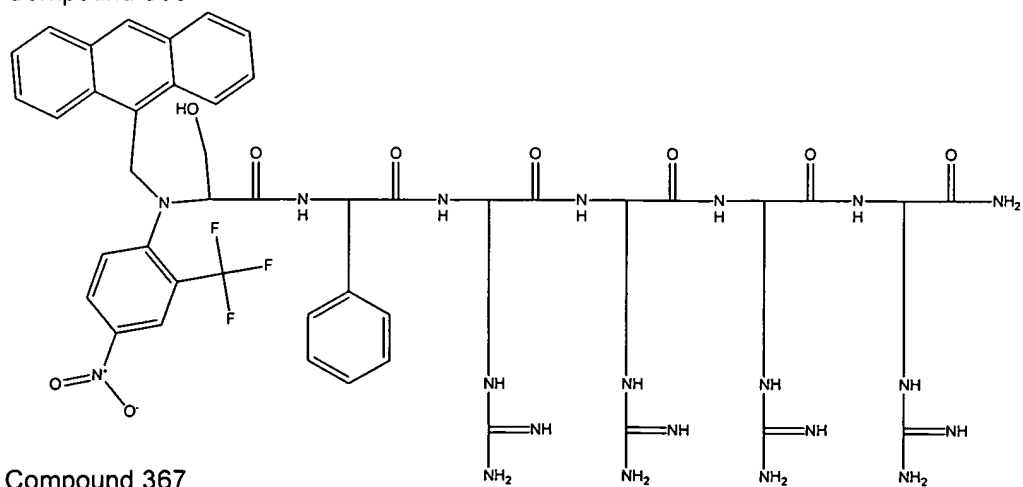
Compound 364



Compound 365



Compound 366



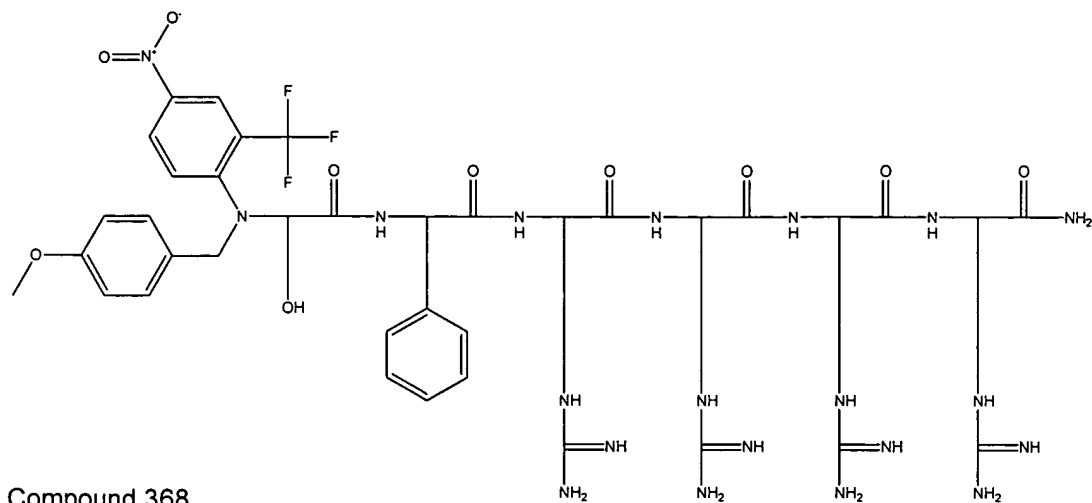
Compound 367

Applicant: David S. Lawrence

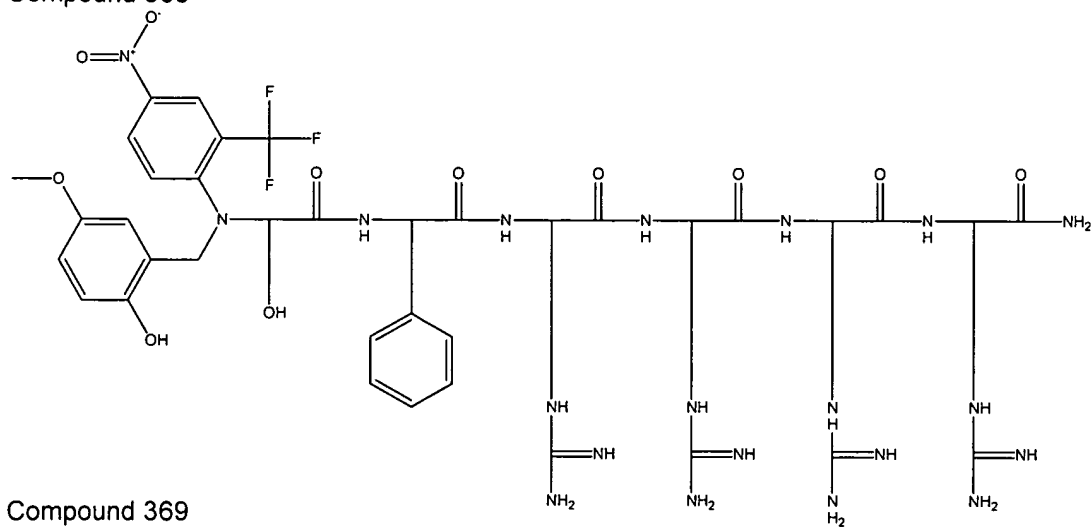
Serial No.: 10/755,086

Filed: January 9, 2004

page 160 of 192



Compound 368



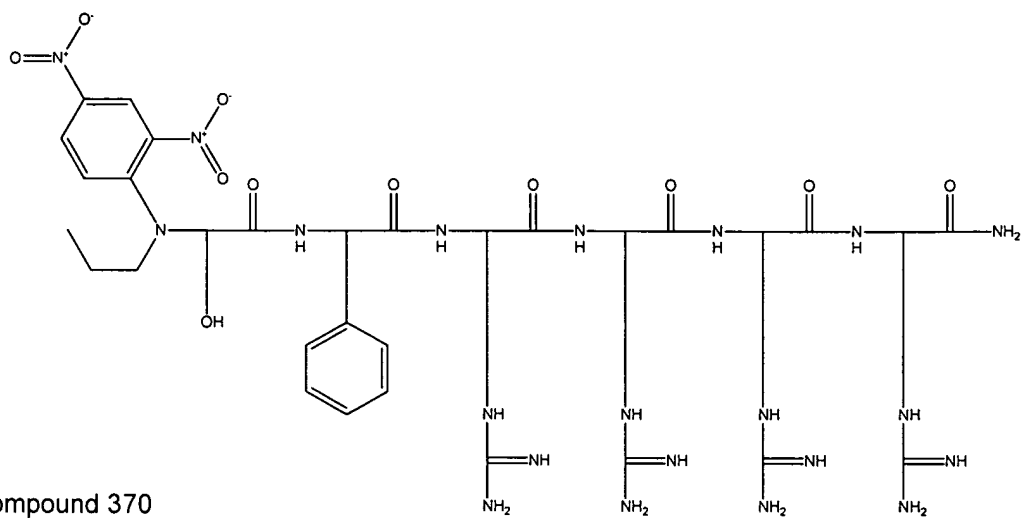
Compound 369

Applicant: David S. Lawrence

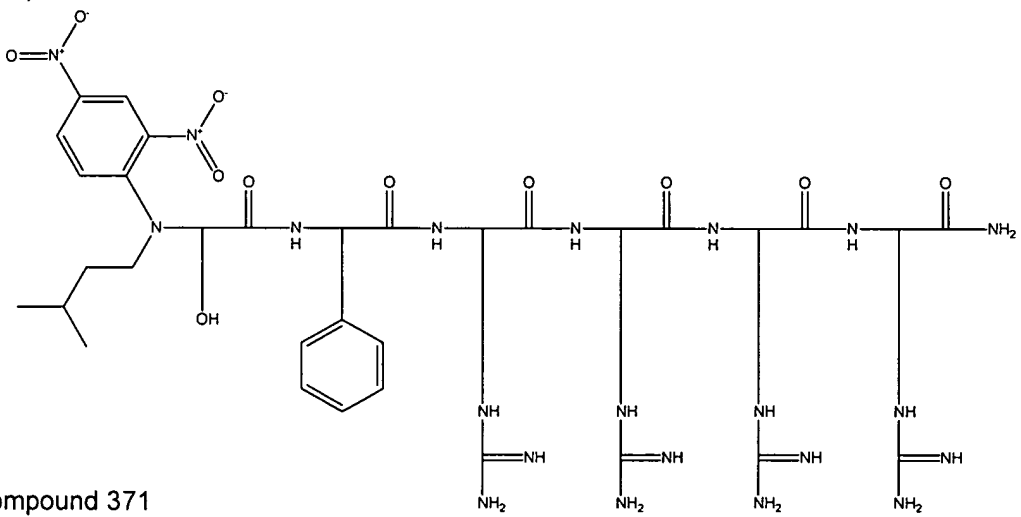
Serial No.: 10/755,086

Filed: January 9, 2004

page 161 of 192

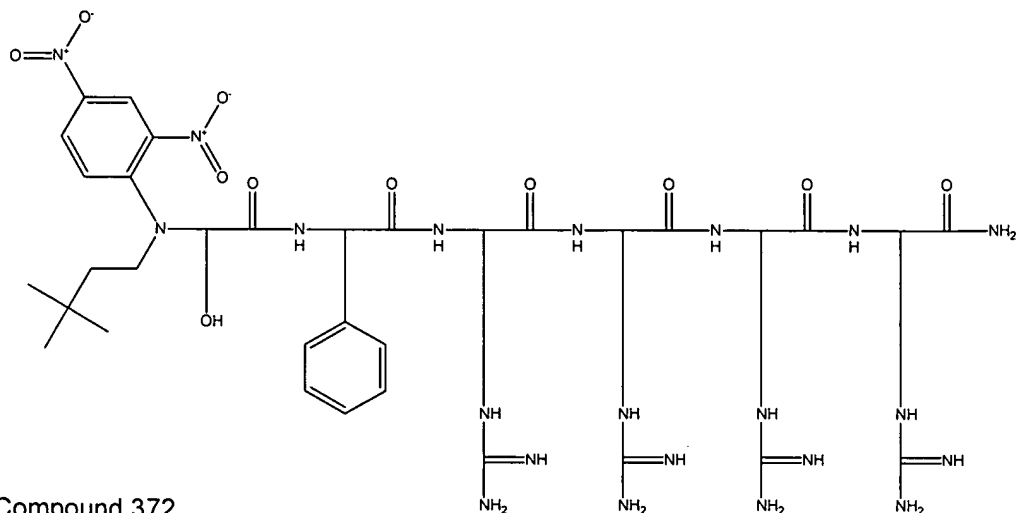


Compound 370

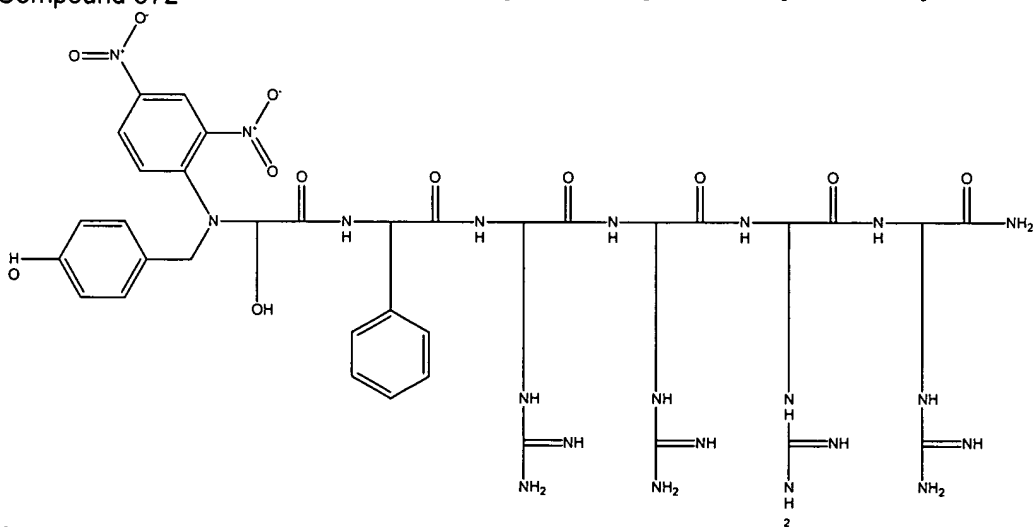


Compound 371

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 162 of 192

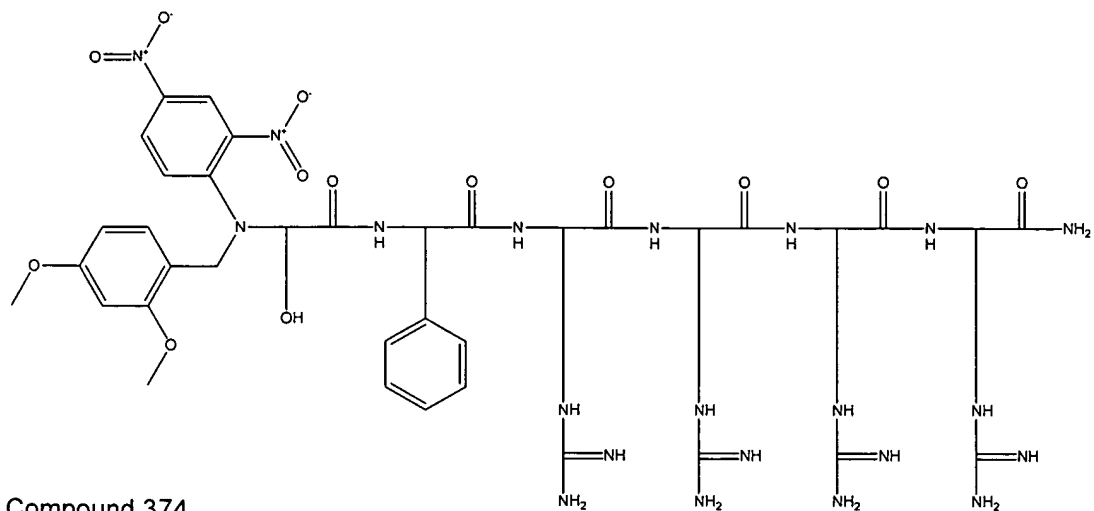


Compound 372

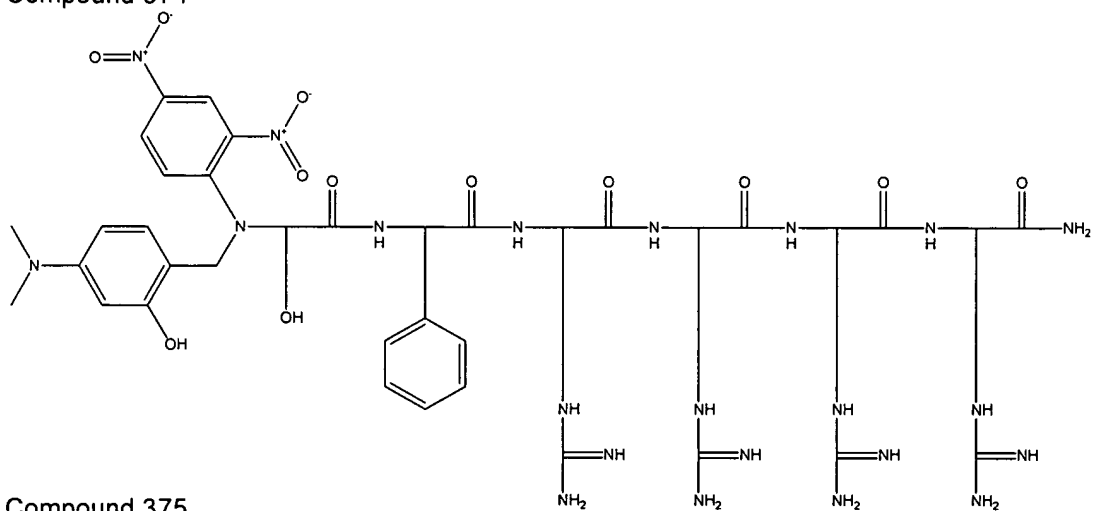


Compound 373

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 163 of 192

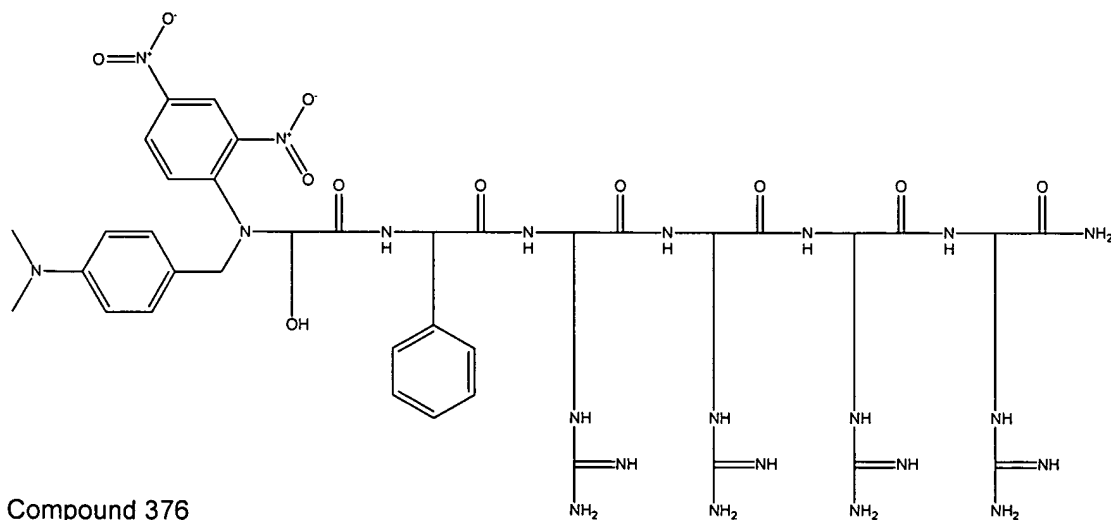


Compound 374

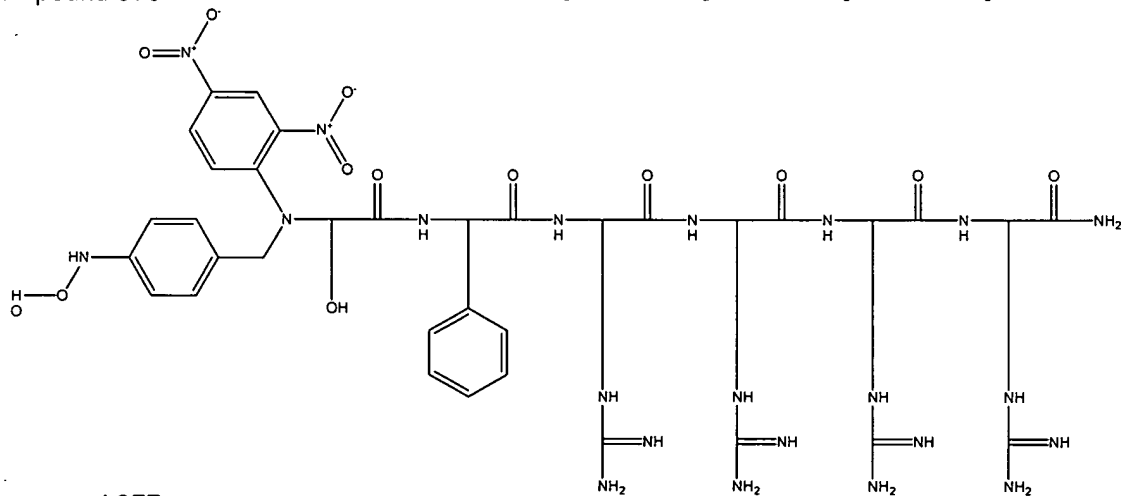


Compound 375

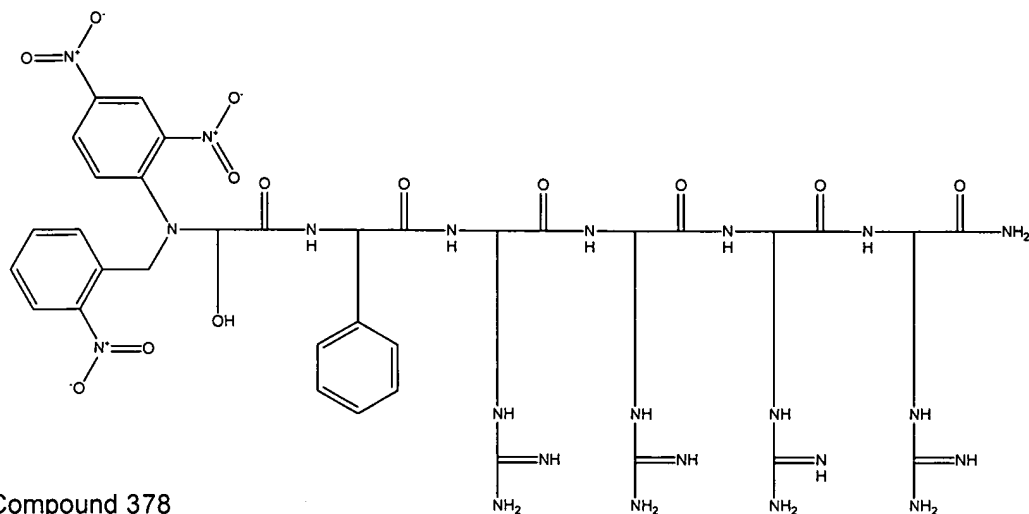
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 164 of 192



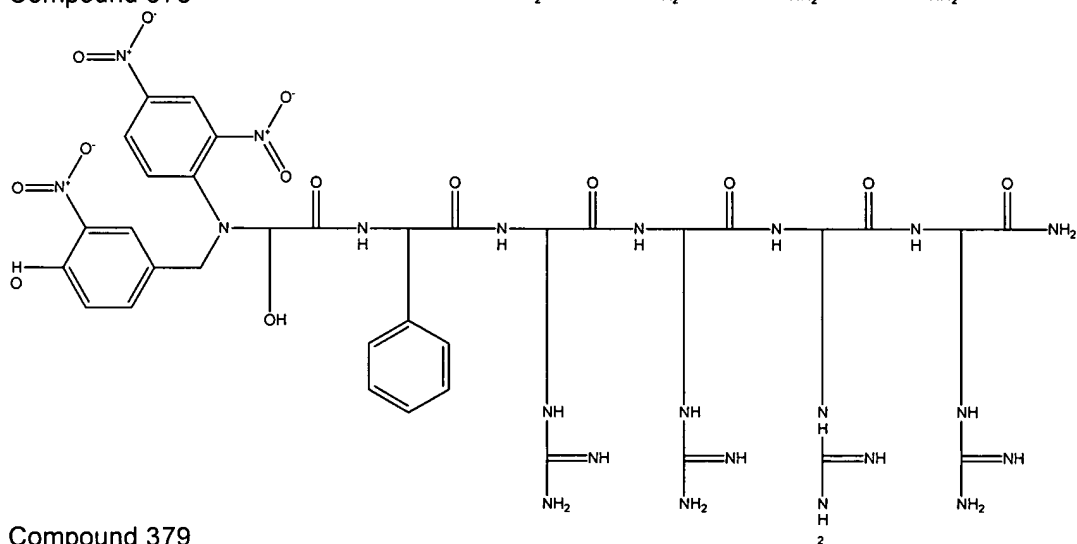
Compound 376



Compound 377

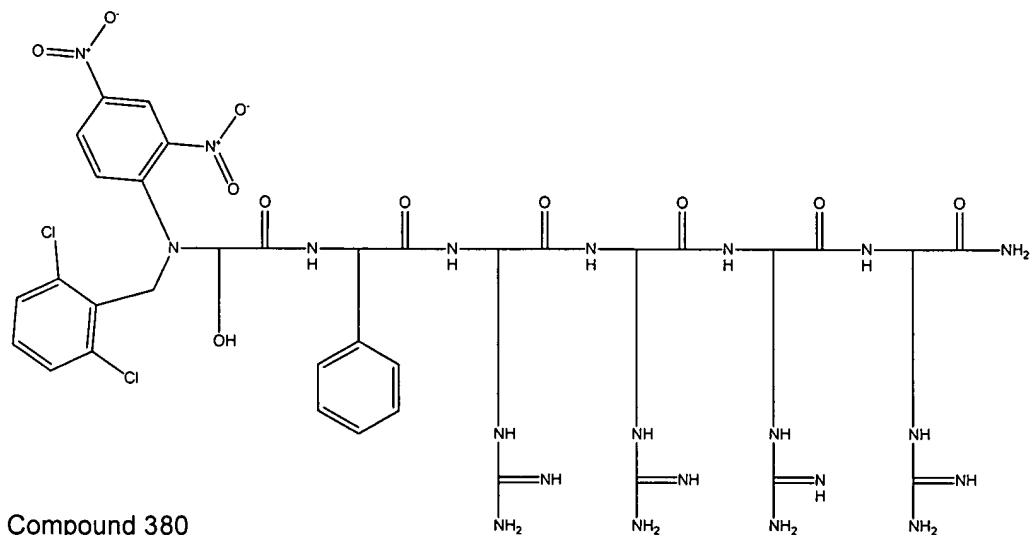


Compound 378

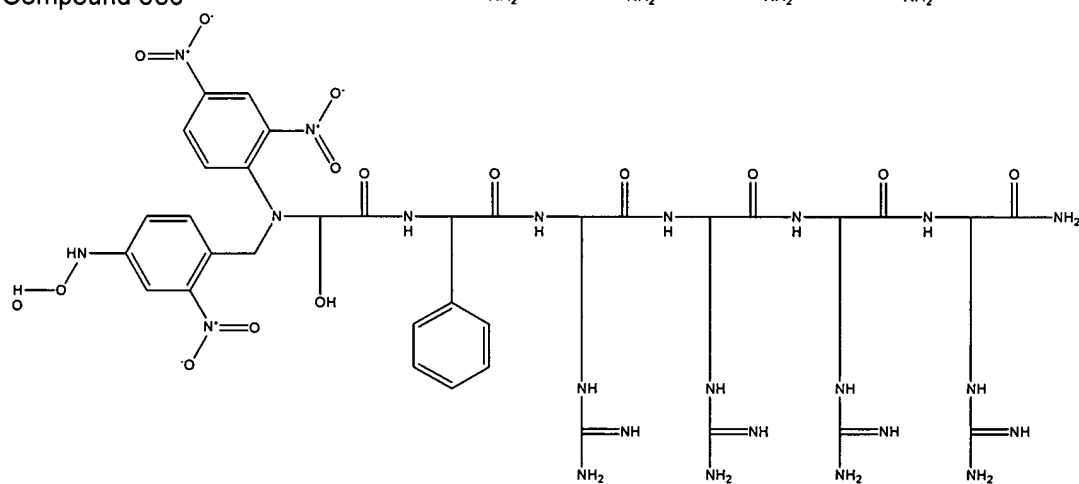


Compound 379

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 166 of 192

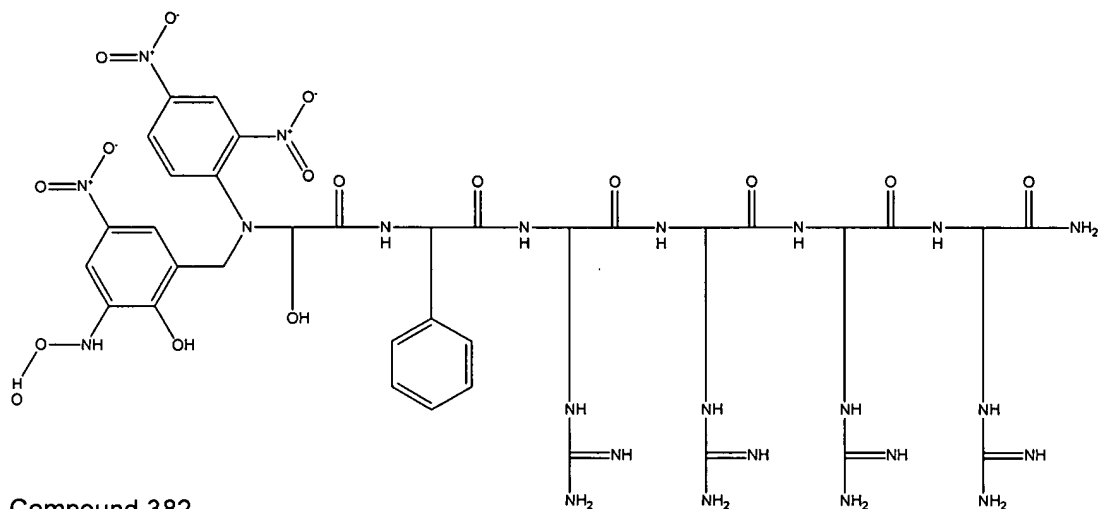


Compound 380

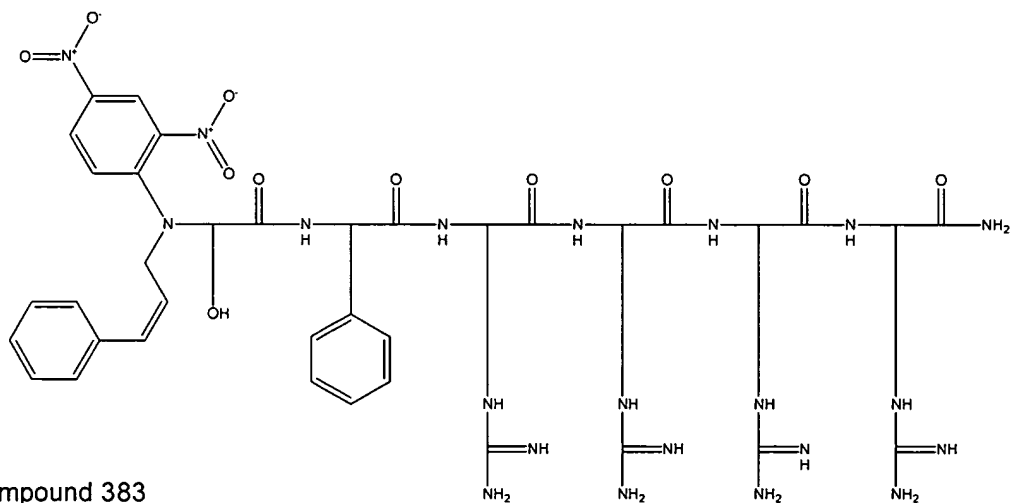


Compound 381

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 167 of 192

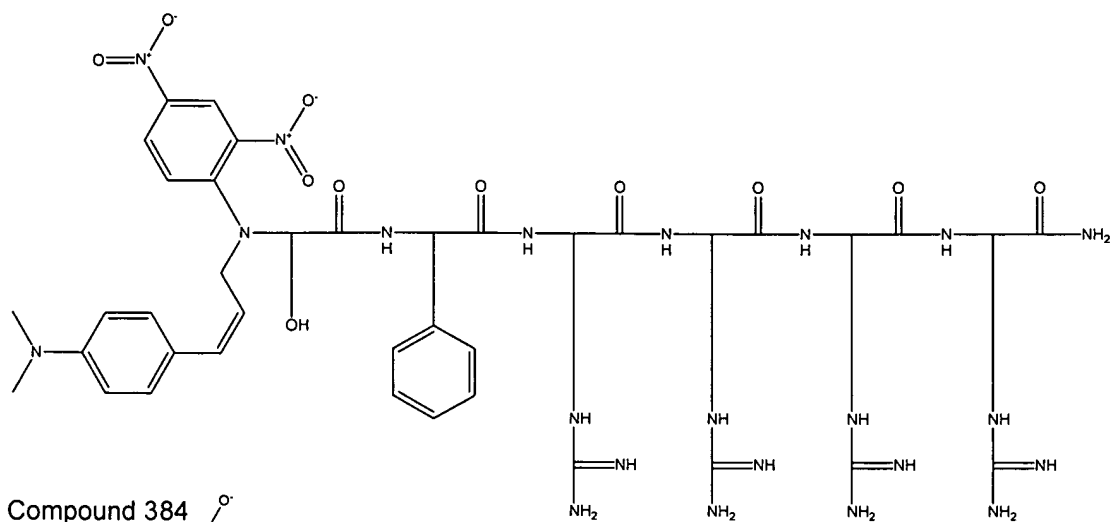


Compound 382

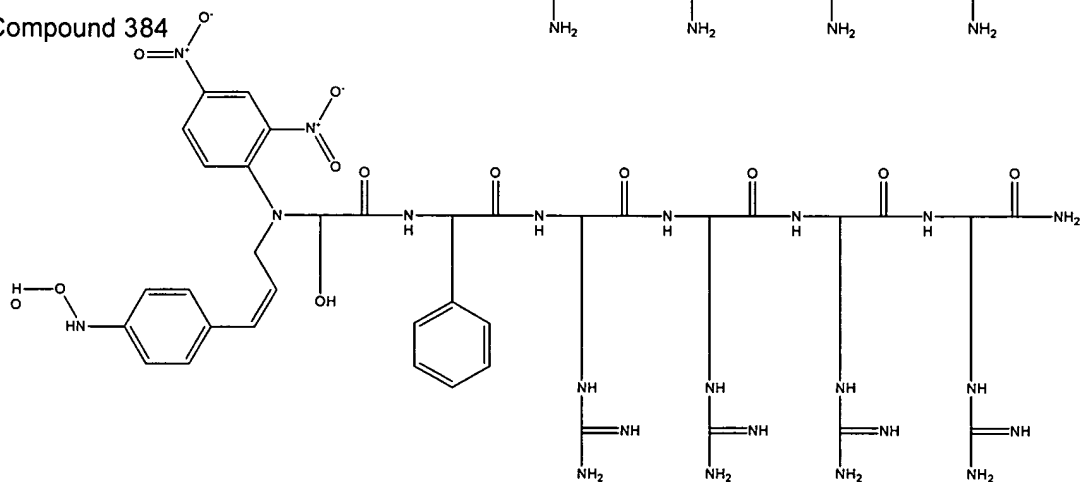


Compound 383

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 168 of 192

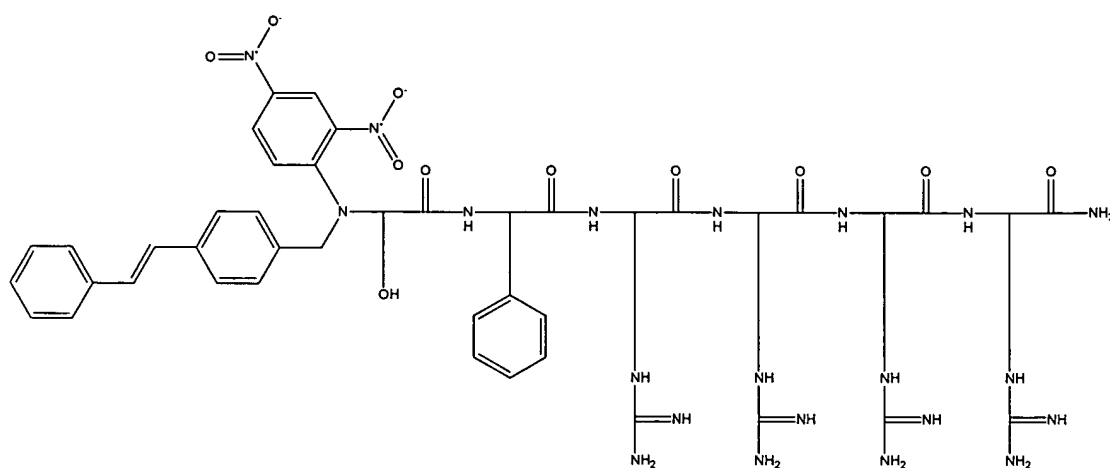


Compound 384

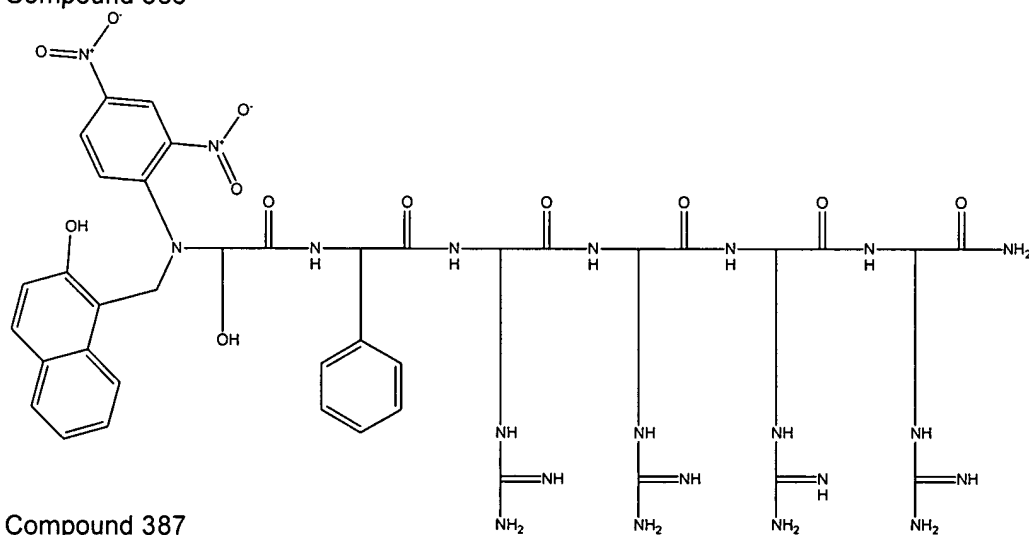


Compound 385

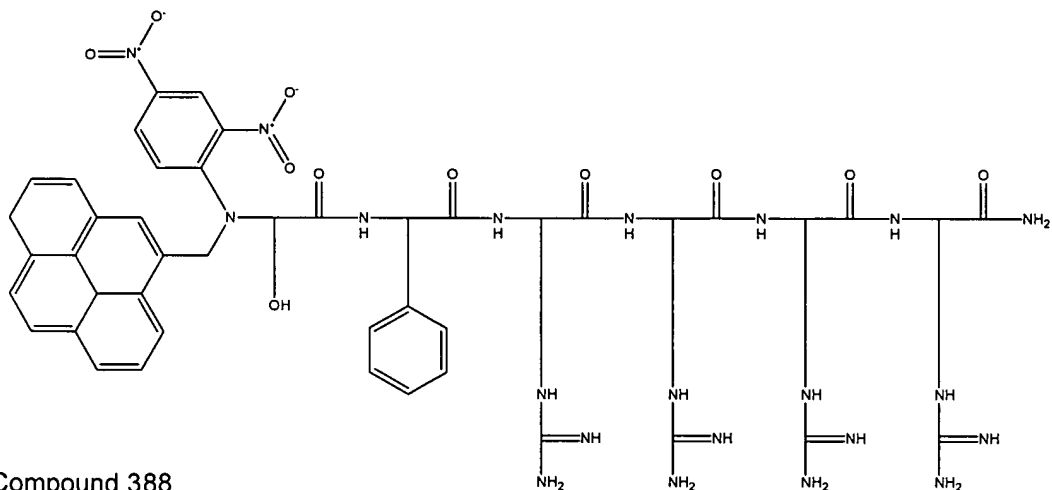
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 169 of 192



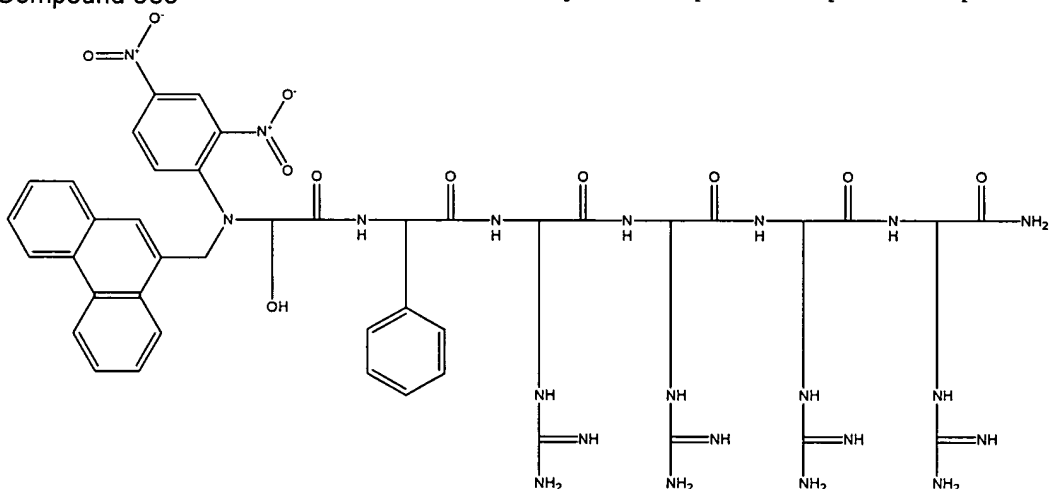
Compound 386



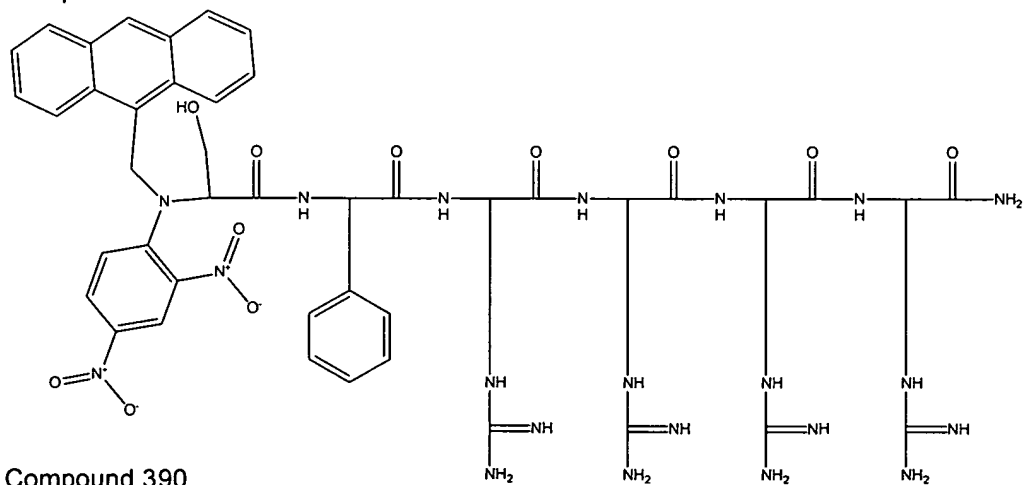
Compound 387



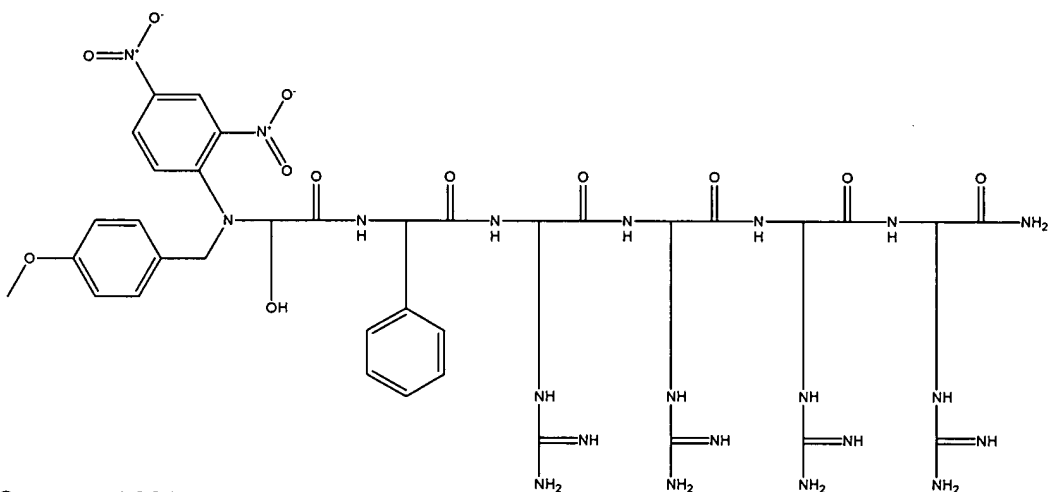
Compound 388



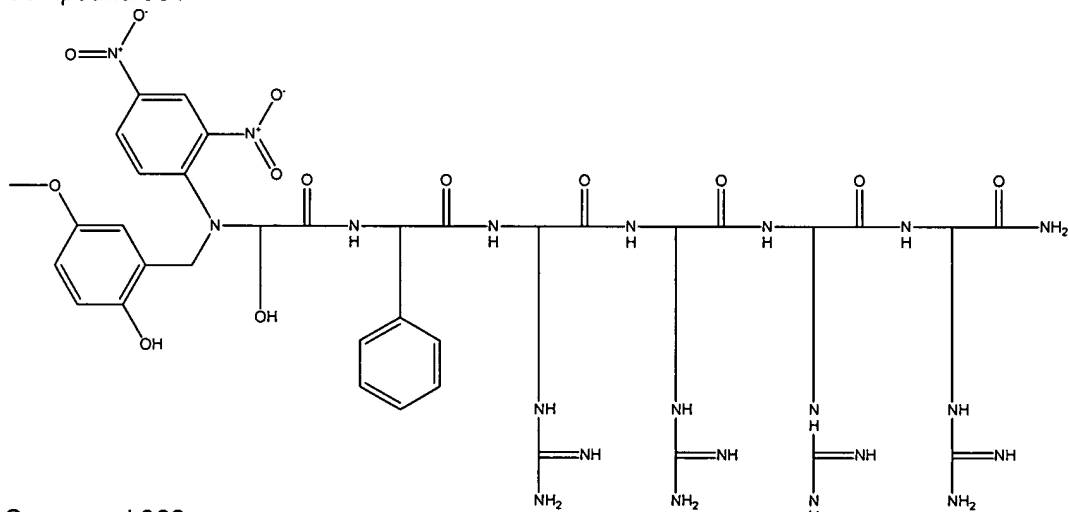
Compound 389



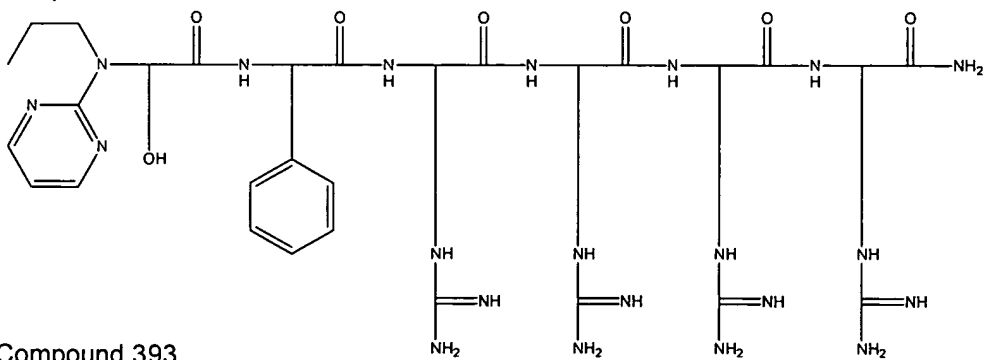
Compound 390



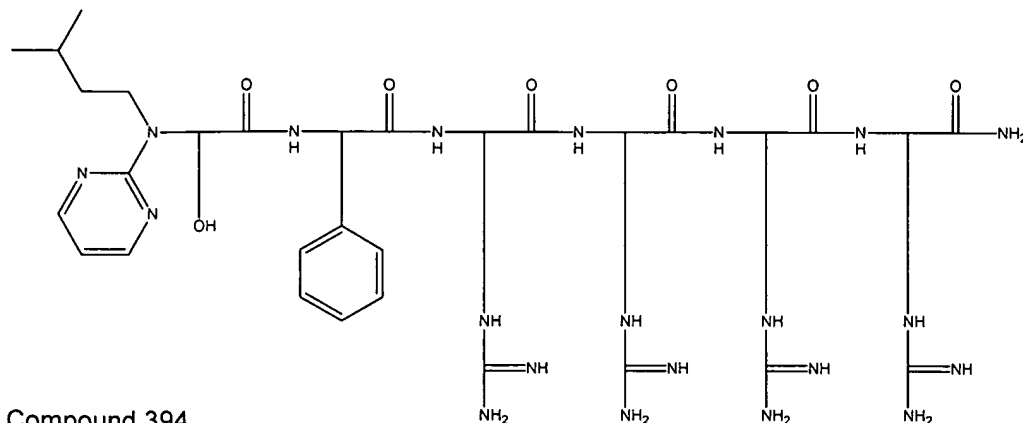
Compound 391



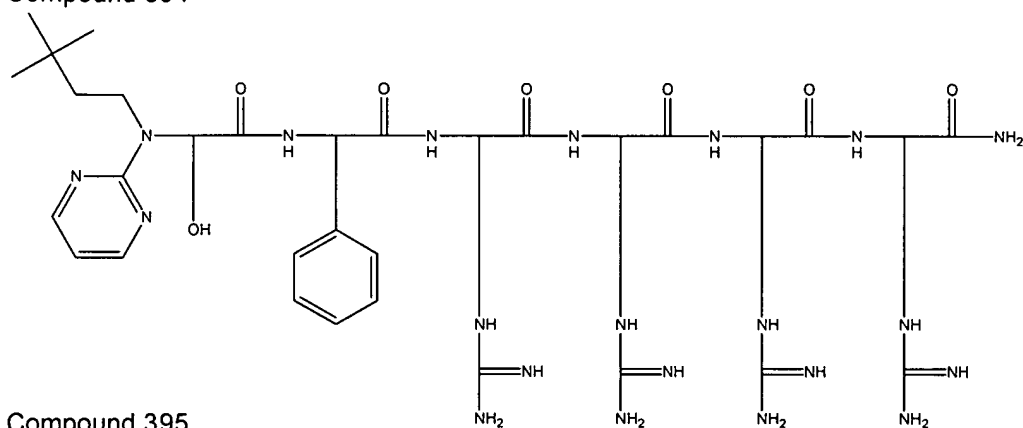
Compound 392



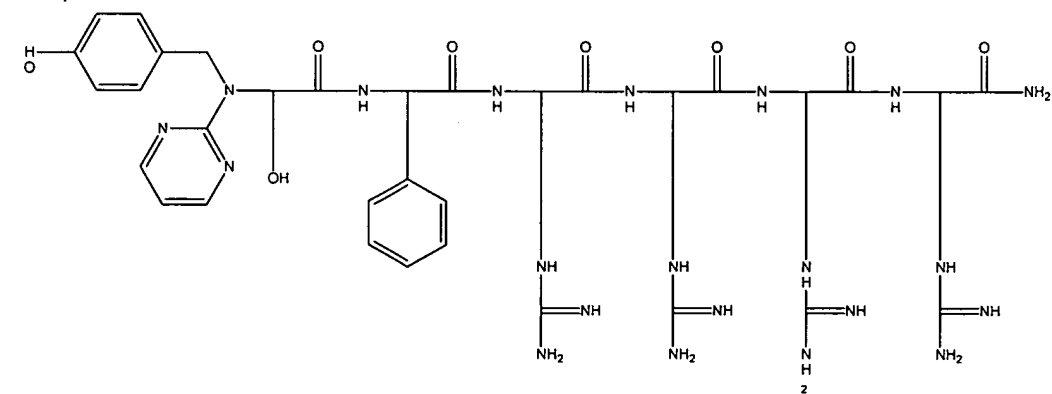
Compound 393



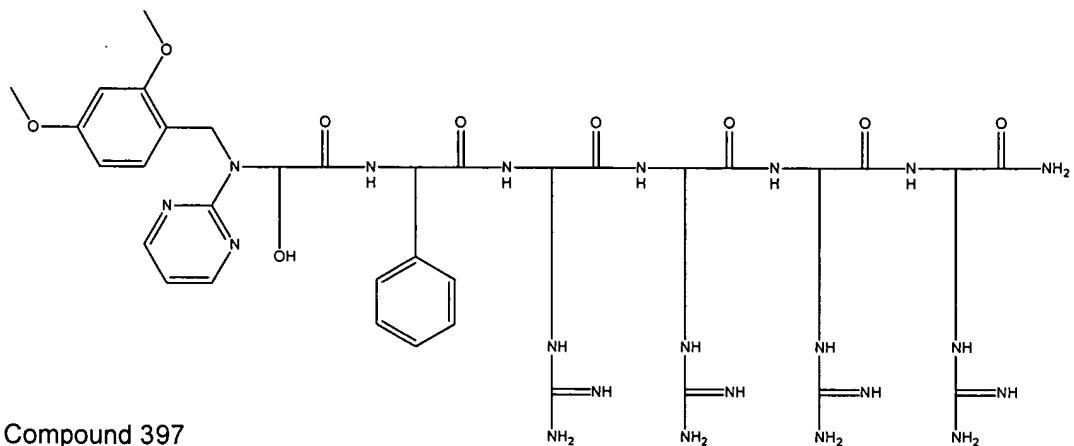
Compound 394



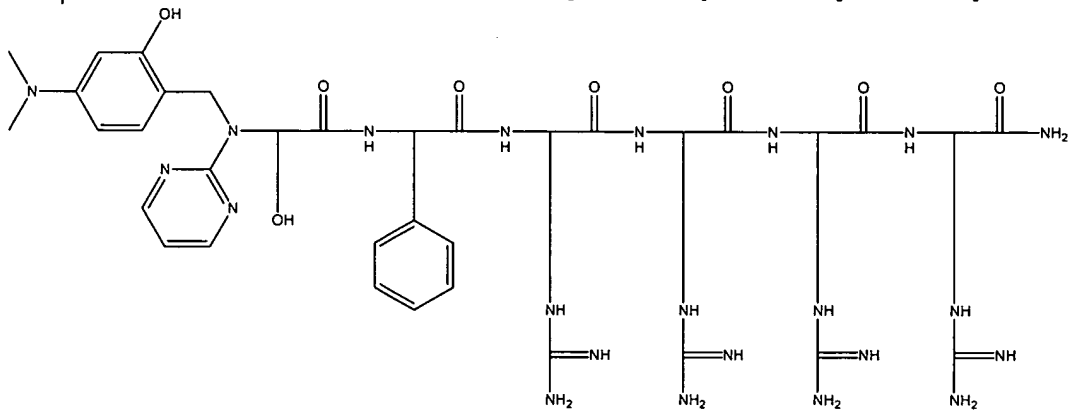
Compound 395



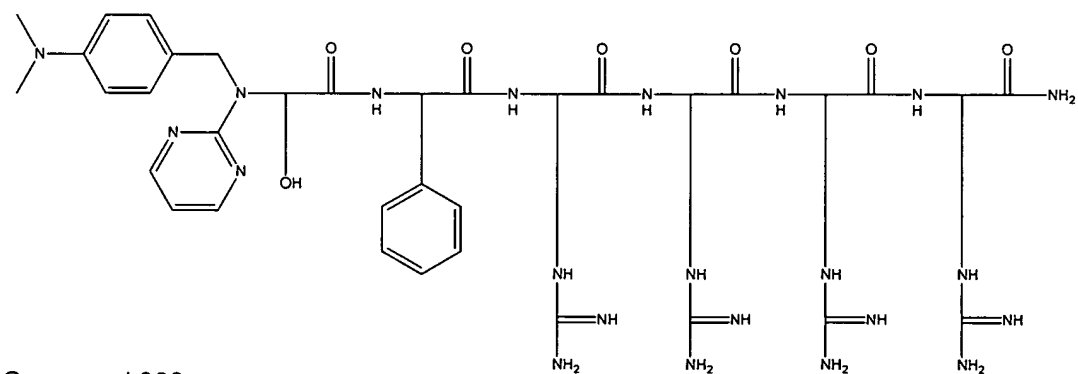
Compound 396



Compound 397

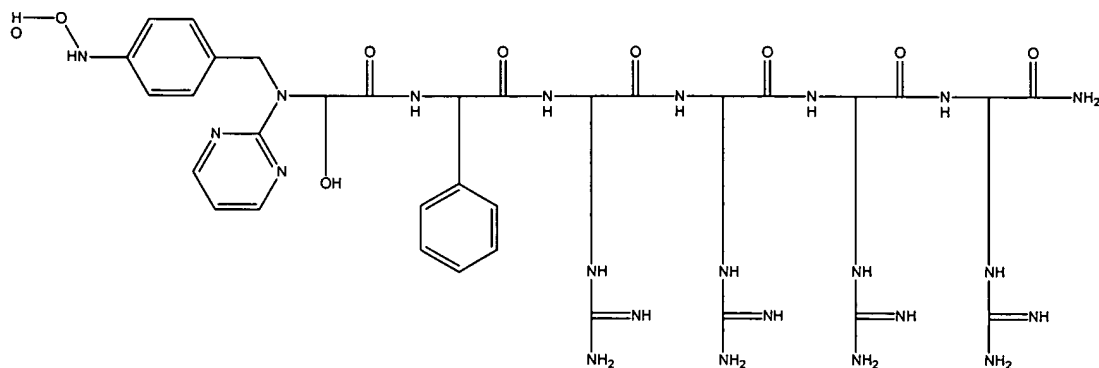


Compound 398

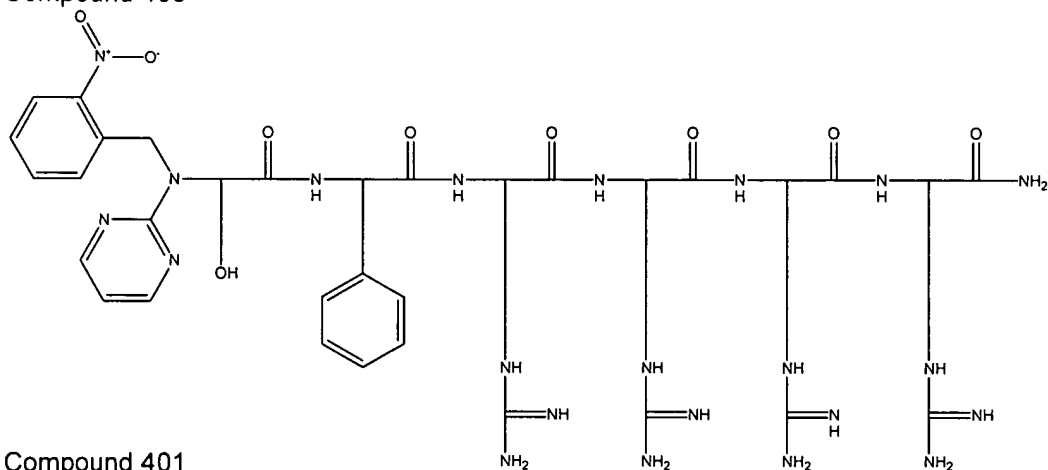


Compound 399

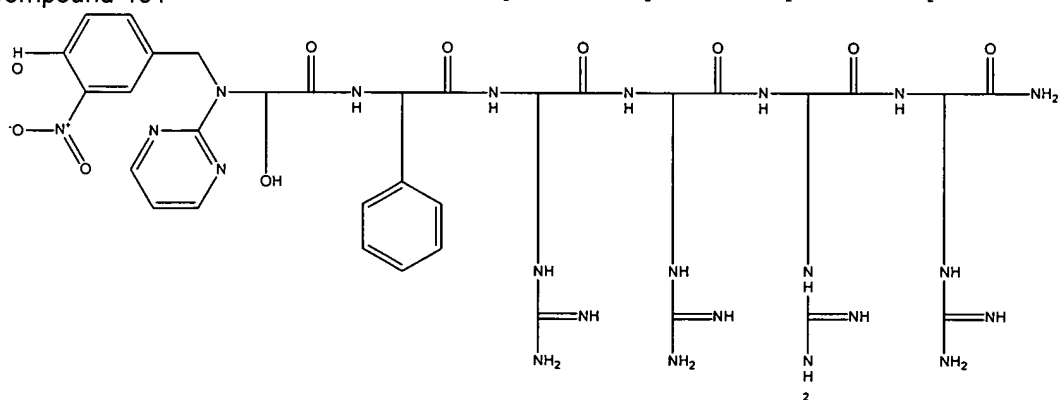
Applicant: David S. Lawrence
 Serial No.: 10/755,086
 Filed: January 9, 2004
 page 174 of 192



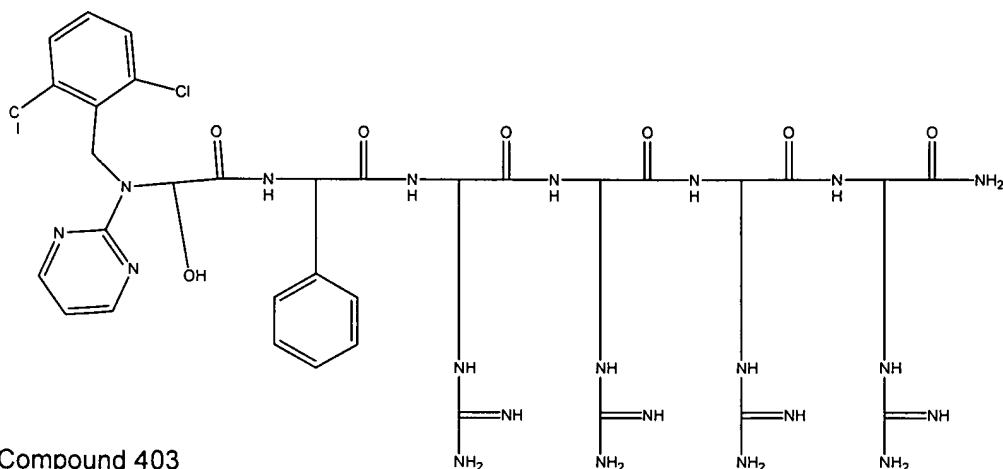
Compound 400



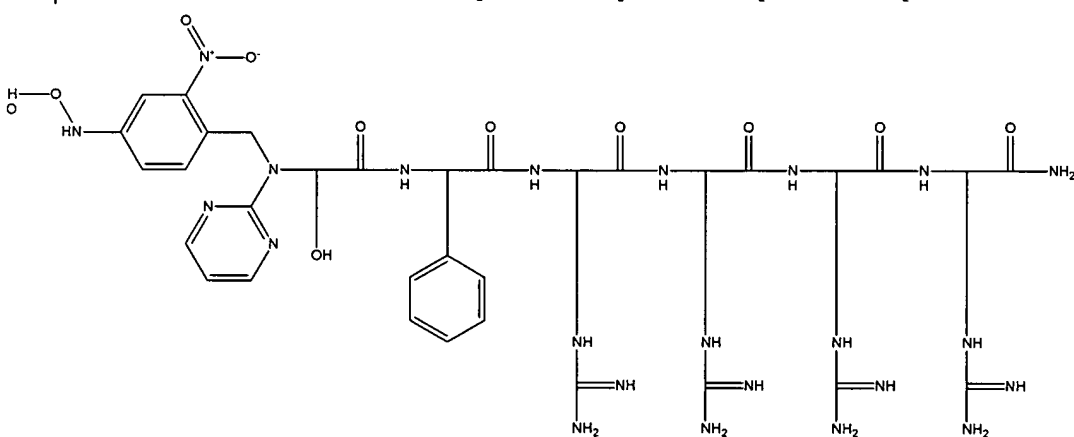
Compound 401



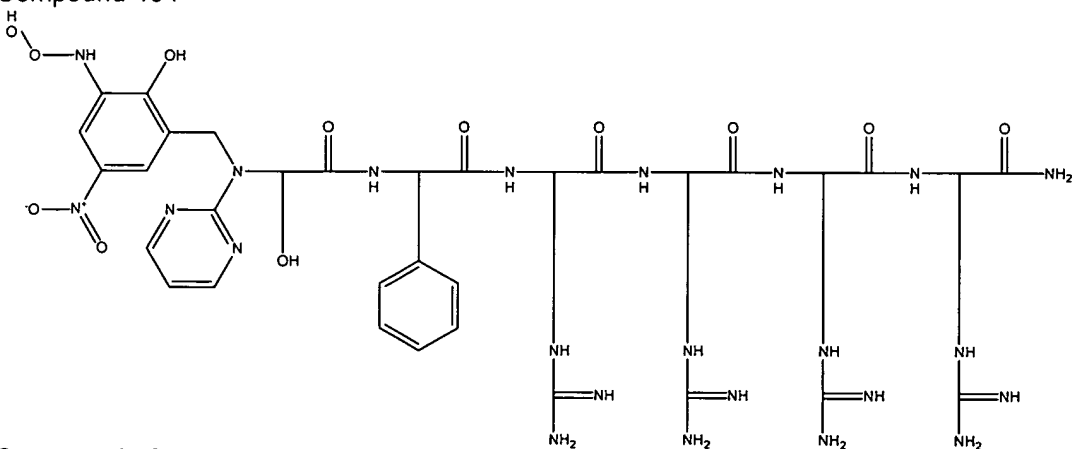
Compound 402



Compound 403



Compound 404



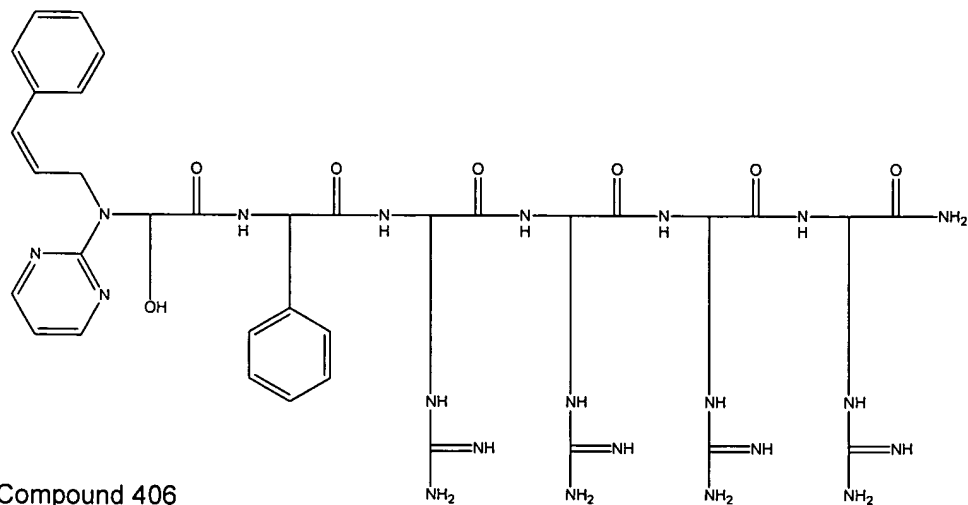
Compound 405

Applicant: David S. Lawrence

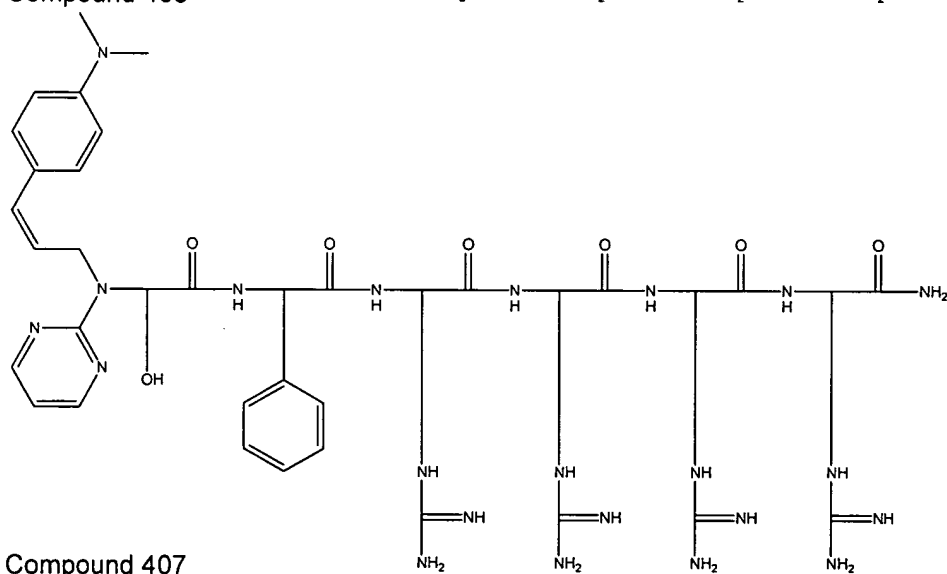
Serial No.: 10/755,086

Filed: January 9, 2004

page 176 of 192



Compound 406



Compound 407

page 177 of 192

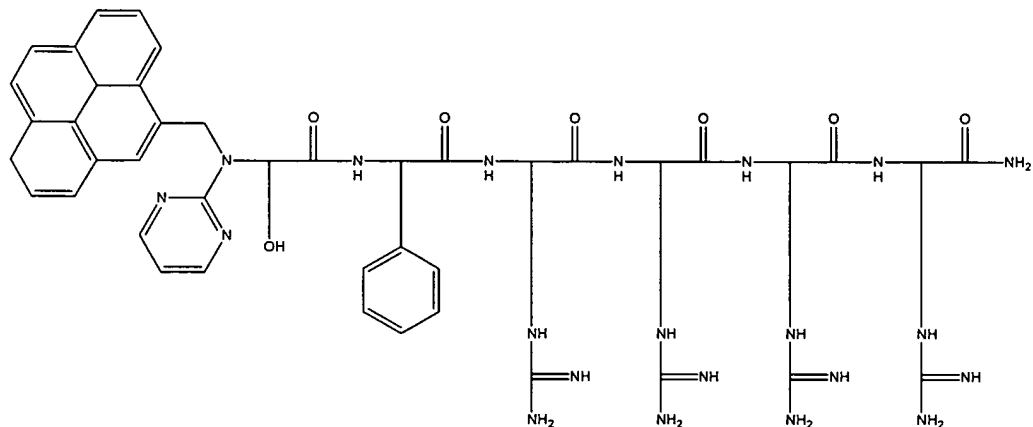


Applicant: David S. Lawrence

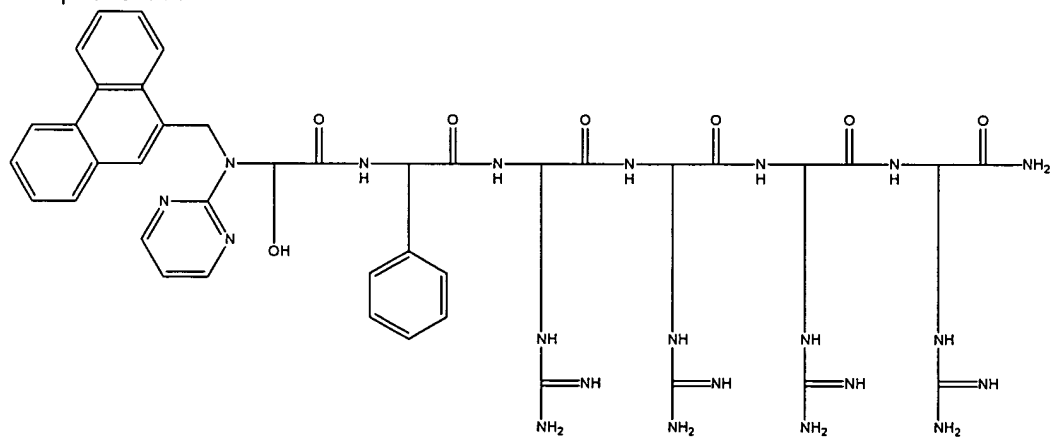
Serial No.: 10/755,086

Filed: January 9, 2004

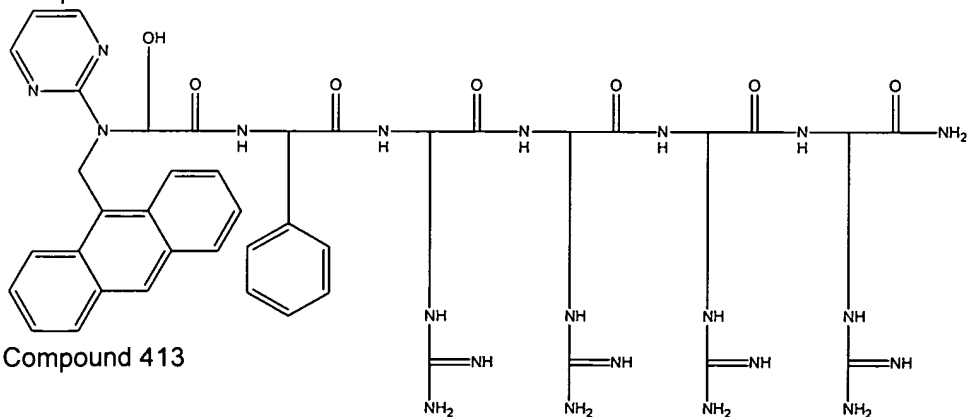
page 178 of 192



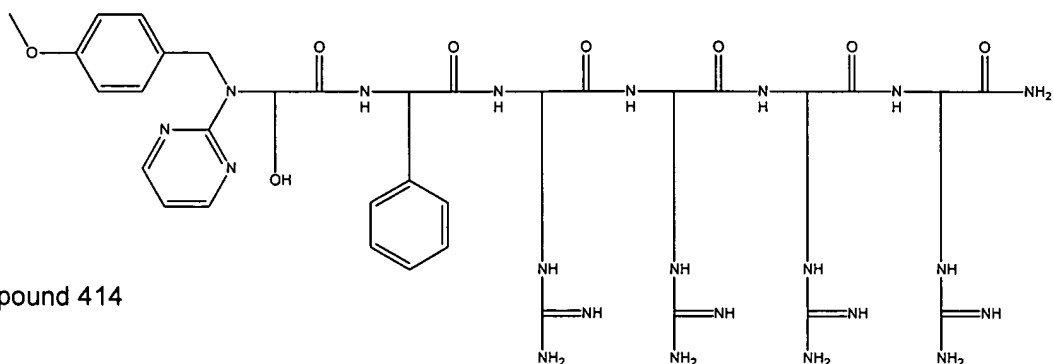
Compound 411



Compound 412

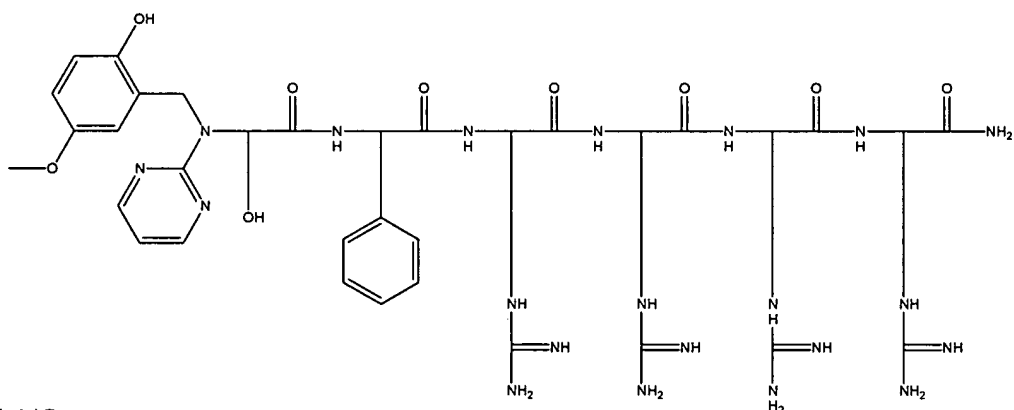


Compound 413



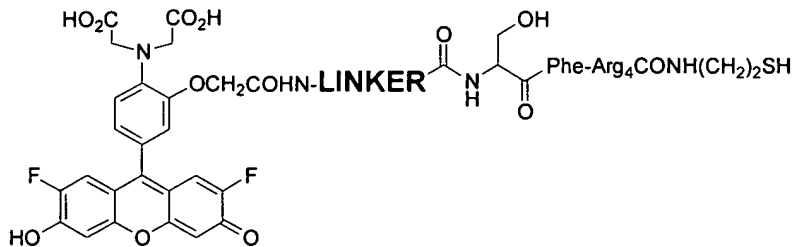
Compound 414

and

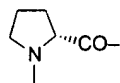


Compound 415

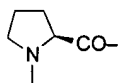
89. (Original) A chemical compound having the structure:



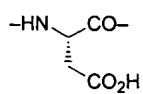
wherein the LINKER is selected from the group consisting of the following:



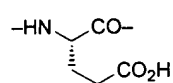
a



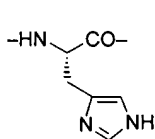
b



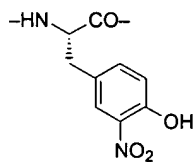
c



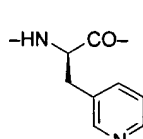
d



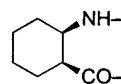
e



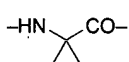
f



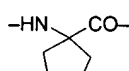
g



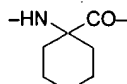
h



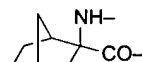
i



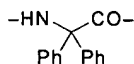
j



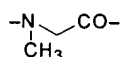
k



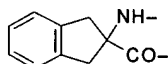
l



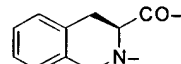
m



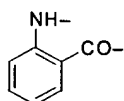
n



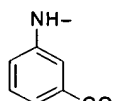
o



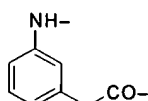
p



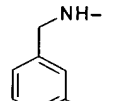
q



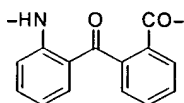
r



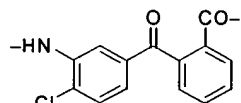
s



t



u



v

no linker

w

90. (Original) A chemical compound having the structure:

fluorophore-LINKER-X-FRRRRK-amide (SEQ ID NO:3);

wherein F is phenylalanine; K is lysine; R is arginine; and X is serine, threonine, or tyrosine.

91. (Original) The chemical compound of claim 90, wherein the fluorophore is a 7-nitrobenz-2-oxa-1,3-diazole derivative.

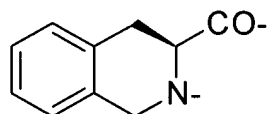
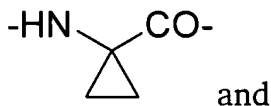
92. (Withdrawn) The chemical compound of claim 90, wherein the fluorophore is a fluorescein derivative.

93. (Withdrawn) The chemical compound of claim 90, wherein the fluorophore is selected from the group consisting of a dansyl derivative, an acridine derivative, an Alexa Fluor derivative, a BODIPY derivative, an Oregon Green derivative, a Rhodamine Green derivative, a Rhodamine Red-X derivative, a Texas Red derivative, a Cascade Blue derivative, a Cascade Yellow derivative, a Marina Blue derivative, a Pacific Blue derivative, an AMCA-X derivative, and a coumarin derivative.

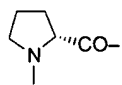
94. (Withdrawn) The chemical compound of claim 90, wherein the linker is a metal chelating linker.

95. (Original) The chemical compound of claim 90, wherein the linker is selected from the group consisting of a carboxamide linker, an aminobenzoic acid linker, a sulfonamide linker, a urea linker, a thiourea linker, an ester linker, a thioester linker, an alkylamine linker, an arylamine linker, an ether linker, and a thioether linker.

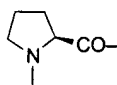
96. (Withdrawn) The chemical compound of claim 90, wherein the linker is selected from the group consisting of N-methyl glycine, L-proline, D-proline,



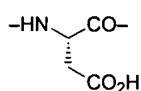
97. (Original) The chemical compound of claim 90, wherein the linker is selected from the group consisting of the following:



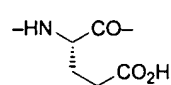
a



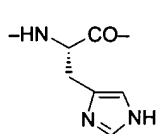
b



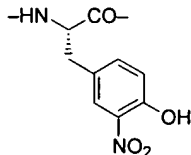
c



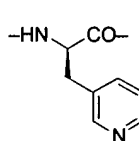
d



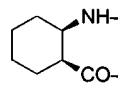
e



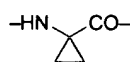
f



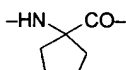
g



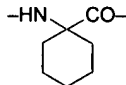
h



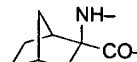
i



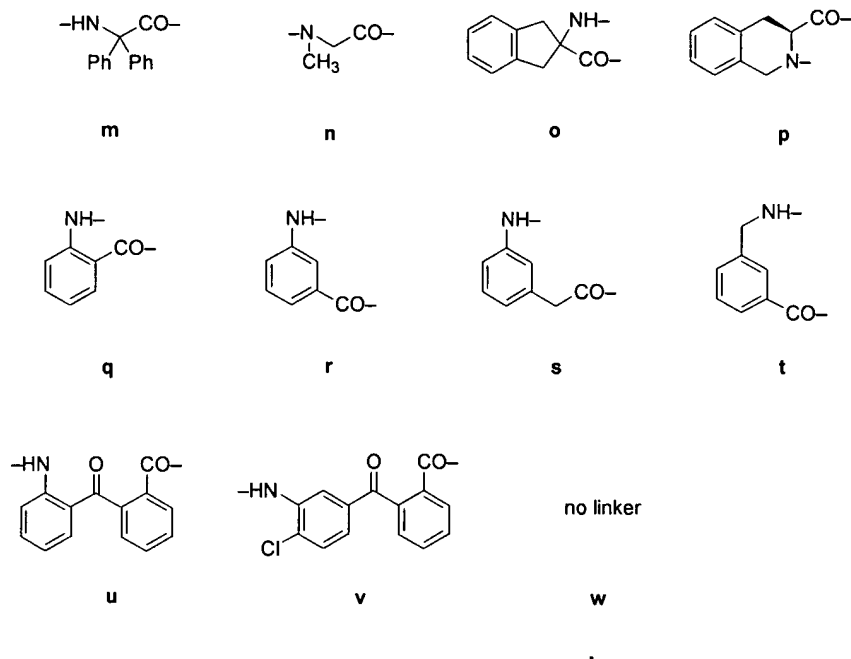
j



k



l



98. (Original) The chemical compound of claim 90, wherein the chemical compound is a substrate for a protein kinase.

99. (Original) The chemical compound of claim 98, wherein the chemical compound is specific for protein kinase C.

100. (Original) The chemical compound of claim 99, wherein the chemical compound is specific for isoforms α , β , and γ of protein kinase C.

101. (Withdrawn) The chemical compound of claim 98, the chemical compound is specific for protein kinase A, protein kinase B, protein kinase D, protein kinase G, Ca^{2+} /calmodulin-dependent protein kinase, mitogen-activated protein kinase, protein kinase mos, protein kinase raf, protein tyrosine kinase, tyrosine kinase abl, tyrosine kinase

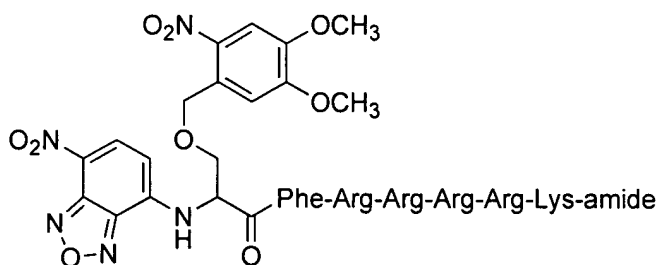
Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 184 of 192

src, tyrosine kinase yes, tyrosine kinase fps, tyrosine kinase met, cyclin-dependent protein kinase, or cdc2 kinase.

102. (Original) The chemical compound of claim 90, wherein the chemical compound further comprises a carbohydrate, a lipid or a nucleic acid.

103-121. (Canceled)

122. (Original) A chemical compound having the structure



123. (Previously presented) A composition comprising a chemical compound of claim 89, and a carrier.

124-126. (Canceled)

127. (Previously presented) The substrate of claim 60, wherein the substrate comprises a metal ion chelator.

128. (Original) The substrate of claim 127, wherein the metal ion is a magnesium ion or a calcium ion.

129. (Currently amended) The chemical compound of claim 90, ~~94~~, wherein a metal ion chelator induces a change in fluorescence intensity.

130. (Original) The chemical compound of claim 129, wherein the metal ion is a magnesium ion or a calcium ion.

131. (Original) The chemical compound of claim 129, wherein the change in fluorescence intensity is at least a 20% change in fluorescence intensity.

132. (Canceled)

133. (Currently amended) The chemical compound of claim 90, ~~substrate of claim 81~~, wherein the linker comprises a turn to position the fluorophore in a location closer to the ~~terminal~~ serine, the ~~terminal~~ threonine or the ~~terminal~~ tyrosine than the location the fluorophore would occupy in the absence of a turn in the linker.

134. (Previously presented) The chemical compound of claim 89, wherein the linker comprises a turn to position the fluorophore in a location closer to the terminal serine, the terminal threonine or the terminal tyrosine than the location the fluorophore would occupy in the absence of a turn in the linker.

135-136. (Canceled)

137. (Previously presented) The composition of claim 123, wherein the composition is a pharmaceutical composition and the carrier is a pharmaceutically

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 186 of 192

acceptable carrier.

138. (Previously presented) A composition comprising the substrate of claim 49, and a carrier.

139. (Previously presented) The composition of claim 138, wherein the composition is a pharmaceutical composition and the carrier is a pharmaceutically acceptable carrier.

140. (Previously presented) A composition comprising the compound of claim 88, and a carrier.

141. (Previously presented) The composition of claim 140, wherein the composition is a pharmaceutical composition and the carrier is a pharmaceutically acceptable carrier.

142. (Previously presented) A composition comprising the compound of claim 90, and a carrier.

143. (Previously presented) The composition of claim 142, wherein the composition is a pharmaceutical composition and the carrier is a pharmaceutically acceptable carrier.

144-145. (Canceled)

146. (Previously presented) A composition comprising the compound of claim

Applicant: David S. Lawrence
Serial No.: 10/755,086
Filed: January 9, 2004
page 187 of 192

122, and a carrier.

147. (Previously presented) The composition of claim 146, wherein the composition is a pharmaceutical composition and the carrier is a pharmaceutically acceptable carrier.